Different target-discrimination times can be followed by the same saccade-initiation timing in different stimulus conditions during visual searches

Tomohiro Tanaka, Satoshi Nishida, and Tadashi Ogawa

1Department of Integrative Brain Science, Graduate School of Medicine, Kyoto University, Kyoto, Japan; and 2Center for Enhancing Next Generation Research, Kyoto University, Kyoto, Japan

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The lateral intraparietal area (LIP) plays roles in visual-motor transformation (Bisley and Goldberg 2010; Gold and Shadlen 2007; Gottlieb and Snyder 2010) and is crucial for visual searching (Liu et al. 2010; Wardak et al. 2002). Changes in the saccadic reaction time in a given stimulus condition could be accounted for by comparable changes in the target-discrimination time in LIP activity, suggesting that the postdiscrimination interval is constant across trials within a condition. Differences in the degree of search speed between different stimulus conditions could be accounted for by comparable changes in the target-discrimination time in LIP activity, suggesting that the postdiscrimination interval is constant across trials within a condition.

Address for reprint requests and other correspondence: T. Ogawa, Center for Enhancing Next Generation Research, Kyoto Univ., Sakyo-ku 606-8302, Kyoto, Japan (e-mail: ogawa.tadashi.7a@kyoto-u.ac.jp).

IN THE REAL WORLD, WE SEARCH for a target object under a variety of stimulus conditions. First, the target is discriminated from distractors using stimulus feature signals; the target spatial information is then linked to the shift in covert attention/overlapping saccades (Schall and Thompson 1999). It is generally thought that the interval required for the former process (prediscrimination interval) varies according to the search difficulty across different stimulus conditions (e.g., Bichot et al. 2001; Sato et al. 2001), whereas the interval for the latter process (postdiscrimination interval) might be stimulus invariant, because stimulus feature signals are no longer necessary after the target location is specified. However, this has not been thoroughly assessed in previous neurophysiological studies.
stimulus presentation, data acquisition, and data analyses have been described elsewhere (Nishida et al. 2013, 2014; Tanaka et al. 2013).

Animal Preparation and Apparatus

Data were collected from two female macaque monkeys (Macaca fuscata; monkey Y, 4 yr old, 5.0 kg; monkey S, 10 yr old, 6.3 kg). Subconjunctival scleral search coils (Judge et al. 1980), head-restraining devices, and recording chambers were implanted during aseptic surgery that was performed under general anesthesia using ketamine and pentobarbital sodium. The recording chambers (22 mm inner diameter) were placed at stereotaxic coordinates (anterioposterior, P0-4 mm; mediolateral, L18-21.5 mm) above the intraparietal sulcus (IPS) guided by T1 magnetic resonance imaging (MRI). All of the animal care procedures and experimental protocols were in accordance with the National Institutes of Health Guidelines for the Care and Use of Laboratory Animals (1996) and were approved by the Animal Care and Use Committee of Kyoto University.

Visual stimuli were generated by a video-signal generator (ViS-aGe; Cambridge Research Systems, Cambridge, UK) and were presented on a video monitor with a 100 Hz refresh rate and an 800 × 600-pixel resolution (RDF223H; Mitsubishi, Tokyo, Japan). Stimuli were viewed binocularly from a distance of 42 cm in a dark room and subtended a visual angle of 51.5 × 40.0°. Extracellular activity was recorded using an epoxylute-insulated tungsten electrode (Frederick Haer & Co, Bowdoinham, ME) with impedance > 2 MΩ (model IMP-1; Bak Electronics, Germantown, MD), and the signal was stored at a sampling rate of 50 kHz. The eye position was monitored with the scleral search-coil technique (Fuchs and Robinson 1966) (eye-position detector DSC-2001; Datel, Tokyo, Japan), and the signals were analyzed at a 1 kHz resolution.

Behavioral Paradigms

Reaction-time visual search task. Figure 1A illustrates the time sequence for a reaction-time visual search task. In each trial, after sustained fixation within a window (± 1.2–1.8°) for 1,000 or 1,500–2,000 ms, a six-element search array was displayed on an imaginary circle (eccentricity = 8.5°). In the target-present trials, the monkeys were required to make a saccade toward a color singleton target after array presentation in the target-present trials. The target stimulus appeared with equal probability in the receptive field or in the location diametrically opposite the fixation spot; therefore, the target appeared at just two fixed locations in each recording session. Due to this design, the correct response rate was slightly higher than reported previously (e.g., Bichot et al. 2001; Thompson et al. 2005; White and Munoz 2011). When the computer detected a saccadic eye movement, the visual stimuli and fixation spot were extinguished immediately. If...
the monkeys made a single saccade that landed inside a square window (± 3.0 × 3.0°) centered on the target, another fixation point appeared at the target position. The monkey received a juice reward after 600 ms of fixation on this point. Target-absent trials were also interleaved with target-present trials. In the target-absent trials, all six elements had the same stimulus features, and the monkeys received a reward if they maintained their fixation (600–1,800 ms) until the end of the trial. The target-present and target-absent trials were presented pseudorandomly at a ratio of 4:1. Because the reaction-time trials were started without any cue that provided information regarding the stimulus condition or target location, the monkeys did not use any top-down bias for target selection (e.g., Bichot et al. 2005; Bichot and Schall 1999; Buschman and Miller 2007; Ogawa and Komatsu 2004, 2009).

Figure 1B shows the four stimulus conditions used for the target-present trials, which were created by manipulating the target-distractor color similarity and luminance of the array elements (Bright-Easy, Bright-Difficult, Dim-Easy, Dim-Difficult conditions). The luminance of each element was either 10 cd/m² (Bright) or 1 cd/m² (Dim) on a gray background (1 cd/m²). The color of each element was either orange (International Commission on Illumination chromaticity coordinates, x = 0.43, y = 0.47 for monkey Y; x = 0.36, y = 0.54 for monkey S), yellowish-orange (x = 0.41, y = 0.48 for monkey Y; x = 0.42–0.43, y = 0.47 for monkey S), green (x = 0.24, y = 0.41 for monkey Y; x = 0.23, y = 0.37–0.39 for monkey S), or bluish-green (x = 0.23, y = 0.38 for monkey Y; x = 0.22, y = 0.34 for monkey S). The target- and distractor-color pairs were selected so that their colors were either more different (Easy: target/distractor colors orange/green, green/orange, yellowish-orange/bluish-green, or bluish-green/yellowish-orange) or more similar (Difficult: yellowish-orange/orange, orange/yellowish-orange, bluish-green/green, or green/bluish-green) (Fig. 1C). Each color pair was presented with equal probability. The color differences between the target and distractor elements were carefully manipulated so that the two array stimuli of the Dim-Easy and Bright-Difficult conditions evoked similar mean reaction times. Therefore, the degree of search speed was balanced in terms of the saccadic reaction time between the Dim-Easy and Bright-Difficult conditions.

Memory-guided saccade task. The memory-guided saccade task (Hikosaka and Wurtz 1983) was used to assess visual, motor, and delay activity and to estimate the receptive field of the cell being studied. A trial of this task was initiated by the appearance of a central fixation point (yellow circle/square or white square spot, luminance 56 cd/m² or 66 cd/m²), and a single circular stimulus (luminance 200 cd/m²) was flashed for 500–700 ms during fixation. After a delay (900–1,800 ms), the fixation point was removed, and the monkeys were required to make a saccade to the location of the target that flashed previously.

Data Collection

We specified the location of the IPS lateral bank that exhibits visual- and saccade-related responses (Barash et al. 1991a,b; Maimon and Assad 2006; Mountcastle et al. 1975) and regarded a cortical region as the LIP where neurons exhibit robust, spatially tuned responses during the delay period of a memory-guided saccade task (Barash et al. 1991a,b; Colby et al. 1996; Gnadt and Andersen 1988; Shadlen and Newsome 2001). Area 7a is known to play important roles in visual searches (e.g., Constantinidis and Steinmetz 2001). To ensure that the samples measured were in the LIP rather than in area 7a, neurons recorded at a depth shallower than 3 mm from the dura surface were excluded from the present analysis (Andersen et al. 1990; Linden et al. 1999; Gifford and Cohen, 2004). Neurons were typically recorded at a depth >5 mm from the dura surface (80.3%). Postoperative structural MRI images were acquired for both monkeys and were used to determine the x-y coordinates of the recording position.

Once a neuron was isolated during the recording sessions, the location of the receptive field was assessed in the memory-guided saccade task for all neurons studied (Tanaka et al. 2013). It should be noted that, because the target-location eccentricity was fixed at 8.5° in the present task, this would have induced suboptimal visual stimulation of each neuron (Schall et al. 2007). In previous studies, the target eccentricity was instead changed according to the location of a neuron’s receptive field, and the size of the stimuli was scaled according to that eccentricity to provide equal visibility. Nevertheless, because the primary aim of the current study was to match the mean reaction times between the Bright-Difficult and Dim-Easy conditions, the use of a fixed eccentricity was a time-saving and pragmatic method for this purpose, and this also excluded the possibility that the difficulty of the task under these two conditions would be imbalanced when the eccentricity was changed (e.g., Gruber et al. 2014; Shive and Francis 2013).

Data Analysis

Unless indicated otherwise, only data from the successful trials were analyzed in this study.

Spike Density Function

The spike train was sampled at 1 kHz and was convolved with a Gaussian of sigma 10 ms to produce a spike density function (Richmond and Optican 1987). We calculated the normalized spike density function for each neuron by transforming the range between the baseline activity during a 200 ms interval before stimulus onset and maximum activity across stimulus conditions during the interval from stimulus onset to the mean saccadic reaction time in the range of 0–1. We constructed the normalized population responses by averaging the normalized spike density functions across neurons.

Target-discrimination Time

The target-discrimination time was defined as the first time point at which target activity (the activity when the target was within a neuron’s receptive field) was significantly greater than distractor activity (the activity when the target was opposite the receptive field) for at least 10 ms (Wilcoxon rank-sum tests, P < 0.01). The length of time from stimulus onset to neuronal target discrimination was defined as the prediscrimination interval, and that from target discrimination to saccade onset was defined as the postdiscrimination interval. Neurons that exhibited target discrimination before both the mean reaction time in stimulus-aligned activity and the saccade onset in saccade-aligned activity were analyzed. The target-discrimination time of the normalized population response was defined as the earliest point at which the target and distractor activities differed significantly for at least 10 ms (Wilcoxon rank-sum tests, P < 0.01).

To calculate the target-distractor discrimination probability, we also performed receiver operating characteristic (ROC) analysis in a way that was similar to a method described previously (Thomas and Paré 2007; Thompson et al. 1996). At each millisecond time point, the separation of the distribution of target and distractor activity was quantified by calculating area under the ROC curves (AUCs) as a function of time. Next, the time course of discrimination probability (time sequence of AUCs) was fitted to the cumulative Weibull distribution function in the form W(t) = γ − (γ − δ)exp[−(t/α)β], where t = the time from when the AUC reached its minimum value (δ) to its maximum value (γ), α = the time at which the AUC reached the minimum area under the ROC curve reached the sum of 63.2% of its maximum, and β = the slope. The functions were terminated when the number of either target or distractor trials was <5.
**Goal-discrimination Time in False-alarm Error Trials**

Catch trials occasionally resulted in false-alarm errors in which the monkeys made a saccade to one of the array elements before the trial ended. We compared the activity when the saccade goal was located in the receptive field with the activity when the saccade goal was located diagonally opposite the fixation point from the receptive field. Using an analytic method that was similar to the aforementioned method employed to determine target-discrimination time, we evaluated the goal-discrimination time at which a neuron reliably predicted the goal location of an erroneous saccade response. We analyzed neurons in which the monkey made ≥7 false-alarm saccades to both the receptive field and the location opposite to the receptive field in either the Bright or Dim catch conditions.

**Covariability between the Target-Discrimination Time and Saccadic Reaction Time**

To assess the temporal relationship between the target-discrimination time and the saccadic reaction time, the covariability of these two variables was examined (Thompson et al. 1996). For each neuron, trials under each stimulus condition were divided into two groups according to the saccadic reaction time (short vs. long reaction-time groups), and the target-discrimination time was calculated for each reaction-time group. If there were a tight link between the target-discrimination time and the saccadic reaction time, the changes in the target-discrimination time might be equal to those in the saccadic reaction time, indicating the ratio of a change in the target-discrimination time to a change in the saccadic reaction time was 1. In contrast, if there were no link, changes in the target-discrimination time might be zero regardless of changes in the saccadic reaction time, indicating that the ratio was 0.

**Reaction-time Matching**

The mean saccadic reaction times between the Bright-Difficult and Dim-Easy conditions were balanced by manipulating the properties of the stimulus (Fig. 1). Nevertheless, a small difference in the mean saccadic reaction time and a substantial difference in the variability remained. To eliminate any possible effects of these differences on the present results, we further performed reaction-time matching. A pair of trials from the Bright-Difficult and Dim-Easy trials were selected for each neuron such that the saccadic reaction-time difference did not exceed 5 ms, and this choice was repeated as long as possible. This procedure ensured that the reaction-time distributions were statistically indistinguishable between the two stimulus conditions.

**Leaky Integration of Visually Responsive LIP Neuronal Activities**

Recent studies suggest that visually responsive neurons in the LIP and FEF provide signals that drive an accumulation process in downstream neurons (movement neurons in the FEF and SC), which then initiate a saccade response when their activity reaches a fixed response threshold (Heitz and Schall 2012; Purcell et al. 2010, 2012; Schall et al. 2011). To explore this possibility, the population-averaged raw histograms (1 ms resolution) for the trials when the target appeared in the receptive field were convolved with a leaky integrator, which was modeled as a single exponential (a varying time constant, 1–140 ms) to produce the accumulated responses. The time constant of the leaky integrator was adjusted so that the sum of the differences in the accumulated responses of the four stimulus conditions was minimal at the time of saccade initiation.

**RESULTS**

**Neuronal Database**

The activity of 122 visually responsive neurons in the LIP of two macaques (71 from monkey Y, 51 from monkey S) performing the visual search task was recorded. The neurons that showed target discrimination before the mean reaction times in stimulus-aligned activity and before the saccade onset in saccade-aligned activity were selected and analyzed. Of the 122 cells, 93, 87, 95, and 81 fulfilled this criterion in the Bright-Easy, Bright-Difficult, Dim-Easy, and Dim-Difficult conditions, respectively. The variation in the number of neurons across conditions might be due to differences in the discrimination probability between the Easy (93–95 neurons) and Difficult (81–87 neurons) conditions (Nishida et al. 2013). Of these, 78 fulfilled the criterion in all four stimulus conditions, and 44 (23 from monkey Y and 21 from monkey S) still met the criterion, even when the trials were divided into short and long reaction-time groups in the Bright-Difficult and Dim-Easy conditions. These 44 cells were used for subsequent analyses.

**Behavioral Performance**

Figure 1D shows the saccadic reaction times obtained from the recordings of the 44 neurons studied. The cumulative reaction-time distributions for each neuron were calculated and then averaged across all neurons. The mean reaction time (triangles) was shortest in the Bright-Easy trials (gray) and longest in the Dim-Difficult trials (black). The reaction times largely overlapped and were indistinguishable between the Bright-Difficult (orange) and Dim-Easy (blue) conditions (Wilcoxon signed-rank test, \( P = 0.55 \)), although the variability was larger in the Bright-Difficult condition compared with the Dim-Easy condition. Figure 1E shows the percentage of correct response rates across stimulus conditions. On average, the correct response rates in the Easy conditions (range, 96–97%) were significantly greater than were those in the Difficult conditions (83–85%; Wilcoxon signed-rank test, \( P < 0.001 \)). Thus, although the reaction times were indistinguishable, the correct response rates were significantly different between the Bright-Difficult and Dim-Easy conditions. The possible effects of the differences in the rate of correct responses are considered below.

**Target-discrimination Time under Different Stimulus Conditions**

Figure 2A shows the spike density functions of an example LIP neuron in the Bright-Difficult (Fig. 2A, orange traces, top) and Dim-Easy (Fig. 2A, blue traces, bottom) conditions aligned separately according to stimulus onset (Fig. 2A, left) and saccade onset (Fig. 2A, right). The visually responsive activity was quite different between the two stimulus conditions. The response onset latency (squares), defined as the time point at which the response was greater than the mean + 2 SDs of the baseline activity during a 200 ms period before stimulus onset, was 25 ms earlier in the Bright-Difficult condition (68 ms) than in the Dim-Easy condition (93 ms). In addition, the strength of the initial visual response 70–120 ms after stimulus onset for the Bright-Difficult condition (28 spikes/s) was more than twice that for the Dim-Easy condition (13 spikes/s). In spite of this difference in activity, there was no distinguishable difference in the mean reaction times between conditions (mean ± SE = 200 ± 5.6 ms vs. 199 ± 2.1 ms; two sample t-test, \( P = 0.86 \)).

For the primary purpose of this study, we compared the target-discrimination time (vertical dashed lines) between the
two stimulus conditions. In stimulus-aligned activity, the pre-
discrimination interval was 31 ms shorter in the Dim-Easy
condition (117 ms) compared with the Bright-Difficult condi-
tion (148 ms). Conversely, in saccade-aligned activity, the 
postdiscrimination interval was 36 ms longer in the Dim-Easy 
condition (82 ms) compared with the Bright-Difficult condi-
tion (46 ms). Figure 2B shows the time course of the discrimination 
probability and the best-fit Weibull functions in each condition. The 
discriminability increased earlier and was stronger in the 
Dim-Easy condition compared with the Bright-Difficult condi-
tion. Obvious differences were observed \(~ 100 \text{ ms}\) after the 
stimulus and before the saccade onset.

In addition to the difference in the target-discrimination 
time, we also noticed differences in the response strength at the 
target-discrimination times (horizontal dashed lines in Fig. 2A). 
In both stimulus- and saccade-aligned activity, the instanta-
neous discharge rate at the target-discrimination time was 
\( ~30\% \) stronger in the Bright-Difficult condition (33 spikes/s) 
compared with the Dim-Easy condition (25–26 spikes/s) (Wil-
coxon rank-sum test, \( P < 0.001 \)). Therefore, the results of the 
example neuron suggest that the target-discrimination time and 
response strength at that time could differ among stimulus 
conditions, even if they were followed by similar reaction-time 
saccades.

Population Analyses of Target-discrimination Time under 
Different Stimulus Conditions

The response profiles observed in the example neuron shown 
in Fig. 2 were preserved at the population level. Figure 3A shows 
the normalized population responses under the Bright-
Difficult and Dim-Easy conditions obtained from 44 LIP neu-
rons, which revealed significantly discriminated activity in all four stimulus conditions (Wilcoxon rank-sum test, \( P < 0.01 \)). 
Figure 3, B and C, shows the distributions of the target-
discrimination times and the response strengths at the target-
discrimination times, respectively, obtained from the individ-
ual neurons. The target-discrimination time was significantly 
earlier in the Dim-Easy condition compared with the Bright-
Difficult condition (Wilcoxon signed-rank test, \( P < 0.0001 \)), 
and the response strength at the target-discrimination time was 
significantly stronger in the Bright-Difficult condition than in 
the Dim-Easy condition (Wilcoxon signed-rank test, \( P < 
0.01 \)). Consistent with this, the strength of the target-discrimi-
nation probability increased earlier and was stronger in the 
Dim-Easy condition compared with the Bright-Difficult condi-
tion (Fig. 3D). A significant difference in the discrimination 
probability started \(~ 100 \text{ ms}\) after the stimulus onset and 
remained continuous until the saccade-initiation time (thick 
lines; Wilcoxon signed-rank test, \( P < 0.01 \)).

The results of Fig. 3 were generally consistent across neu-
rons and among monkeys. Figure 4 shows the changes in either 
the target-discrimination time (Fig. 4A) or the response strength at that time (Fig. 4B) for individual neurons between 
the Dim-Easy and Bright-Difficult conditions. Squares and 
circles indicate the data from monkeys \( Y \) and \( S \), respectively. 
The target discrimination occurred earlier in the Dim-Easy 
condition than in the Bright-Difficult condition for 95% (42/
44) of the neurons in stimulus-aligned activity and 84% (37/44) 
of the neurons in saccade-aligned activity. The ratio was 
significantly higher than would be expected by chance in both 
cases \( (\chi^2\text{-test}, P < 0.0001) \). The strength of the response at the 
target-discrimination time was greater in the Bright-Difficult 
condition than in the Dim-Easy condition in 68% (30/44) of 
the neurons in stimulus-aligned activity and in 73% (32/44) of 
the neurons in saccade-aligned activity \( (\chi^2\text{-test}, P < 0.005) \).

Taken together, the population analyses reveal that target 
discrimination occurred earlier in the Dim-Easy condition than 
in the Bright-Difficult condition, whereas the strength of the 
response at the discrimination time was greater in the Bright-

Fig. 2. Activity of a single lateral intraparietal area (LIP) neuron in the 
Bright-Difficult and Dim-Easy conditions. A: spike density functions (mean ± 
SE) in the Bright-Difficult (orange traces) and Dim-Easy conditions (blue 
traces). Left and right panels indicate stimulus- and saccade-aligned activity, 
respectively. Solid and dashed traces of the spike density functions denote the 
activity when the target appeared in the receptive field and away from it, 
respectively. Vertical dashed lines with a number above indicate the target-
discrimination times, and horizontal dashed lines indicate the response strength 
at the target-discrimination time. Gray ticks and black circles above the spike 
density functions indicate the action potentials and saccade onsets, respect-
ively. Each row of rasters indicates 1 trial. Triangles above the raster plots 
indicate the mean saccadic reaction times. B: time-course of the discrimination 
probability at each millisecond as evaluated using receiver operating charac-
teristic (ROC) analysis (circles) and best-fit Weibull functions (thick traces). 
The goodness of fit \( (r^2) \) in the Bright-Difficult and Dim-Easy conditions was 
0.98 and 1.0 for stimulus-aligned activity, and 0.92 and 0.99 for saccade-
aligned activity, respectively. RF, receptive field.
Difficult condition compared with the Dim-Easy condition. In other words, late target discrimination was associated with a strong response (Bright-Difficult), whereas early target discrimination was associated with a weak response (Dim-Easy). The observation that the postdiscrimination interval differed among stimulus conditions suggests that this interval is not devoted solely to processing spatial information for saccade preparation/initiation.

**Reaction-time Matching**

Although the mean reaction times were balanced between the Bright-Difficult and Dim-Easy conditions (Figs. 3, 4), the reaction-time variability was not matched (Fig. 1D). To confirm that the present results were not due to this difference, we performed reaction-time matching between the two stimulus conditions for each neuron to equalize the reaction-time distributions in either the target and distractor trials (AUC = 0.46–0.56; Kolmogorov-Smirnov test, \( P = 0.54–1.0 \)). Figure 5A shows the population mean of the reaction-time distributions across neurons, which were almost perfectly matched between the Bright-Difficult and Dim-Easy conditions (Kolmogorov-Smirnov test, \( P = 1.00 \)). Figure 5, B–E, shows the results when the same analyses used in Fig. 4, A–D, were applied to neuronal data after reaction-time matching. The results described in Fig. 4 remained the same; therefore, the target-discrimination time could differ under different stimulus conditions, even when the reaction times were almost perfectly matched between conditions.

**Which is More Important, Stimulus Luminance or Target-distractor Similarity, for Determining the Postdiscrimination Interval?**

Because both luminance and target-distractor similarity (search difficulty) differed between the Bright-Difficult and Dim-Easy conditions, it was unclear which factor was more important for producing changes in the postdiscrimination interval. To explore this further, we performed two-way ANOVA using neuronal data obtained during all four stimulus conditions (Bright-Easy, Bright-Difficult, Dim-Easy, and Dim-Difficult conditions). Figure 6A shows the target-discrimination time in stimulus-aligned activity. The interval increased with search difficulty and decreased with increasing luminance. Data revealed significant main effects of both target-distractor similarity and luminance [target-distractor similarity, \( \text{F}(1,172) = 158.9, P < 0.0001; \) luminance, \( \text{F}(1,172) = 40.1, P < 0.0001 \)] but no interaction [\( \text{F}(1,172) = 183, P = 0.18 \)]. In contrast, the target-discrimination time in saccade-aligned activity varied only with stimulus luminance (Fig. 6B). Two-way ANOVA showed a significant main effect of luminance [\( \text{F}(1,172) = 17.0, P < 0.0001 \)], but not target-distractor similarity [\( \text{F}(1,172) = 1.35, P = 0.25 \)]; there was no interaction [\( \text{F}(1,172) = 0.01, P = 0.92 \)]. Therefore, the changes at the postdiscrimination interval were likely due to changes at the luminance level, but not in target-distractor similarity (search difficulty).

**Possible Influences of Different Correct Response Rates**

Although the mean saccadic reaction times were balanced between the Bright-Difficult and Dim-Easy conditions (Fig. 1D), a difference in the correct response rates remained (Fig. 1E). Therefore, the reward probability/saccade response probability differed between conditions, and this may have affected...
neuronal activity in the visual-motor areas (e.g., Platt and Glimcher 1999; Sugrue et al. 2004; Thompson et al. 2005). If this were the case, the postdiscrimination interval would vary with search difficulty (i.e., target-distractor color similarity) because the correct response rates significantly varied between the Easy and Difficult conditions (Wilcoxon signed-rank test, \( P < 0.001 \)), but it did not vary with luminance because the correct response rate did not significantly vary between the Bright and Dim conditions (Wilcoxon signed-rank test, \( P > 0.05 \)). However, the present results revealed that the opposite was true (Fig. 6B), indicating that the observed difference in the postdiscrimination interval was likely not explained by a difference in the correct response rate.

**Dependence on History of Search Difficulty**

As shown in Fig. 1D, the correct response rates in the Easy search condition were significantly higher than were those in the Difficult search condition, suggesting that a different degree of search accuracy may be required in the Easy and Difficult search conditions. To ascertain whether the effects of this difference in the speed stress remained in the following trials, we compared the behavioral and neuronal performances in each of the Bright-Difficult and Dim-Easy conditions when the preceding trial was an Easy condition and when it was a Difficult condition. The data were analyzed only when two successive search trials were completed successfully. The results revealed that the saccadic reaction times were statistically indistinguishable (Wilcoxon signed-rank test, \( P = 0.23–0.84 \)), and the strength of the activity did not differ at any millisecond time point 250 ms after stimulus onset (Wilcoxon signed-rank test, \( P > 0.05 \)), irrespective of the search difficulty of the preceding trial. This result suggested that the history of search difficulty did not have a substantial influence on the behavioral and neuronal responses of the following trial.

**Covariability of Target-discrimination Time and Saccade-Initiation Time within Stimulus Conditions**

Previous studies in the LIP revealed that the variability of the saccadic reaction times could be explained by variability in the target-discrimination times (Ipata et al. 2006; Thomas and Paré 2007). To show that the current sample neurons had similar response properties, we examined the relationship between the target-discrimination time and the saccadic reaction times by dividing the trials into two equally sized (short and long reaction-time) groups for each neuron and then calculated the target-discrimination times and the mean reaction times for each group for each neuron. Figure 7A shows the relationship between the target-discrimination time in stimulus-aligned activity (predisiscrimination interval) and the saccadic reaction time obtained under the Bright-Difficult (Fig. 7A, left) and Dim-Easy (Fig. 7A, right) conditions. Each line plot illustrates how a change in target-discrimination time was related to a
change in the mean reaction time for each neuron. Figure 7B shows the slopes of the individual line segments (change in the prediscrimination interval/change in the saccadic reaction time). Their distributions were significantly different from both 0 (Wilcoxon signed-rank test, \( P < 0.001 \)) and 1 (\( P < 0.01 \)) in both conditions. The mean (± SE) slopes were 0.72 ± 0.07 and 0.71 ± 0.13 in the Bright-Difficult and Dim-Easy conditions, respectively, indicating that the variability in the prediscrimination interval partially (70%) accounted for the variability in the reaction times. In contrast, the target-discrimination time in saccade-aligned activity (postdiscrimination interval) covaried less with the saccadic reaction time (Fig. 7C). The slope values were significantly smaller than 1 under both conditions (Wilcoxon signed-rank test, \( P < 0.001 \)), and there was no systematic difference from 0 [marginally significant under the Dim-Easy condition (\( P = 0.024 \)), but this was not significant under the Bright-Difficult condition (\( P = 0.17 \))]. The mean slopes were −0.21 ± 0.18 and −0.21 ± 0.14 in the Bright-Difficult and Dim-Easy conditions, respectively (Fig. 7D).

The above results reveal that a major part of the reaction-time variability could be explained by variability in the prediscrimination interval. This is roughly consistent with previous reports (Ipata et al. 2006; Thomas and Paré 2007). However, the reduced covariability between the prediscrimination interval and the saccadic reaction time observed in the current study (mean slope = 0.7) compared with previous studies (≈1) might be due to differences in the current task requirement. Although multiple saccade responses were allowed before
foveating the target in the previous studies, only a single saccade was allowed in the current study. Such a difference might affect the preservation of the postdiscrimination interval constancy (Shen and Paré 2011).

Applicability of the Leaky Integration Model

Different luminance levels evoke different magnitude visual responses in LIP neurons (Tanaka et al. 2013), and our control analysis suggested that the postdiscrimination interval decreased with increasing luminance (Fig. 6B). These findings raised the possibility that different magnitude responses were transformed into different postdiscrimination intervals. First, a possible mechanism for this effect was the accumulation-to-threshold (rise-to-threshold) mechanism (Carpenter and Williams 1995; Cisek 2006; Hanes and Schall 1996; Mazurek et al. 2003; Ratcliff et al. 2003; Reddi et al. 2003; Smith and Ratcliff 2004). This model was recently used (Heitz and Schall 2012; Purcell et al. 2010, 2012; Schall et al. 2011) to explain how the activity of visually responsive neurons in the LIP, FEF, and SC was transformed into the activity of movement neurons in the
FEF, SC, and the brain stem, which reaches a fixed threshold level of activation at the time of saccade initiation (e.g., Hanes and Schall 1996; Paré and Hanes 2003; but see Everling et al. 1999, 2000; Jantz et al. 2013). Second, one might argue that another mechanism is also possible. The fact that the raw activity in the four stimulus conditions indistinguishably converged into a common activation level at around 80 ms before saccade (the mean strength was 0.49 at 79 ms before saccade, arrow in Fig. 8A) suggests that the time when the activity achieved this activation level might determine the saccadic reaction time. However, the observation that the response strength of the stimulus-aligned activity in the Bright-Difficult condition (in Fig. 3A, left) exceeded this activation level (= 0.49) at 119 ms after stimulus onset, occurring far before the target-discrimination time (154 ms), is difficult to reconcile with the second hypothesis (Schall et al. 2011). Therefore, we preferred the first hypothesis.

To address the hypothesis that LIP activity is accumulated and used to drive the movement-related neurons in the downstream areas (such as the FEF and/or SC), we initially showed that the accumulation-to-threshold model could explain the saccade-initiation timing, irrespective of stimulus conditions by examining whether the accumulated responses of the LIP samples could reach a common threshold level at saccade onset under different stimulus conditions. Figure 8A shows the raw population activities under the four stimulus conditions, which did not converge into a common activation level at saccade onset (one-way repeated-measures ANOVA, P < 0.01). However, the accumulated responses produced by applying a leaky integration function (time constant = 114 ms) to the raw population activities reached a common threshold level at the saccade onset (one-way repeated-measures ANOVA, P > 0.05) (Fig. 8B).

Next, we show that this accumulation-to-threshold model could explain the dependence of the postdiscrimination interval on stimulus luminance. The accumulated responses calculated from the reaction-time matched trials (Fig. 5) were compared between the Bright-Difficult and Dim-Easy conditions (Fig. 8C). The time constant was set to the optimal value found in Fig. 8B (i.e., 114 ms). The advantage of using reaction-time matched data was that we were able to compare the accumulated responses throughout the trials in the stimulus-aligned format. This allowed us to confirm that the accumulated responses did not exceed the threshold level until saccade onset even when the accumulated responses were aligned to stimulus onset. The result shows that although the target-discrimination times (circles on the accumulated responses) were substantially different (33 ms, 157 vs. 124 ms) between the Bright-Difficult and Dim-Easy conditions, the difference in the threshold-crossing time (i.e., the predicted saccade-initiation timing) was small (4 ms, 231 vs. 235 ms).

Our computer simulation might explain why the postdiscrimination interval differed among conditions. Importantly, the strength of the response at the target-discrimination time differed substantially between the two conditions: the response strength was lower in the Dim-Easy condition (normalized response strength = 0.19) compared with the Bright-Difficult condition (0.48). Therefore, the remaining height, which was increased to reach the threshold, was greater in the Dim-Easy condition than in the Bright-Easy condition. This difference in the response strength meant that it took a longer time for the

![Fig. 8. Applicability of a leaky-integrator model. A: normalized population activity aligned according to saccade onset in the 44 neurons. The 4 traces indicate the spike density functions in the 4 stimulus conditions. Horizontal thick bars indicate the target-discrimination times shown in Fig. 5B. The raw spike-timing histogram (1 ms resolution) aligned according to saccade onset for each neuron was convolved with a leaky integrator and averaged across neurons. An optimal time constant for leaky integration was determined (114 ms) such that the summed difference between each pair of accumulated responses was minimal at the time of saccade initiation. Horizontal gray lines indicate the threshold level (the mean strength of the accumulated responses at saccade onset). B: accumulated responses aligned according to saccade onset. The raw spike-timing histogram (1 ms resolution) aligned according to saccade onset for each neuron was convolved with a leaky integrator and averaged across neurons. An optimal time constant for leaky integration was determined (114 ms) such that the summed difference between each pair of accumulated responses was minimal at the time of saccade initiation. Horizontal gray lines indicate the threshold level (the mean strength of the accumulated responses at saccade onset). C: stimulus-aligned accumulated responses, which were derived from the reaction-time matched Bright-Difficult and Dim-Easy trials (Fig. 5). Because the reaction times were matched between conditions the 2 accumulated responses were compared from the stimulus to the saccade onset. The time constant was set as in B. Solid and dashed traces indicate the accumulated responses when the target appeared in and away from the receptive field, respectively. Target-discrimination times in the accumulated responses (circles on the traces) were the same as the target-discrimination times shown in Fig. 5B. Horizontal gray lines indicate the same threshold level shown in B. Triangles indicate the threshold-crossing times. The increasing accumulated response for each condition was evaluated by linear fitting 160–230 ms after stimulus onset (thick line segments).](http://jn.physiology.org/doi/10.1152/jn.00043.2015)
accumulated response to reach the threshold level after the target-discrimination time in the Dim-Easy condition compared with the Bright-Difficult condition, although the accumulated response rose faster in the Dim-Easy condition (rising rate = 5.2 units/s) than in the Bright-Hard condition (3.7 units/s). Thus, the results of the current simulation suggest that the difference in the strength of the response at the target-discrimination time is an important factor that determines the length of the postdiscrimination interval.

**Target-discrimination Time is Independent of Neuron Type**

Visually responsive LIP neurons can be classified into sub-populations according to the existence of saccade-burst activity or delay activity. To determine whether the target-discrimination time was different depending on the type of neuron, we calculated the target-discrimination time separately for the different types of neurons (Table 1). For 35 of the 44 neurons being studied, the datasets obtained from the memory-guided saccade task were large enough (≥5 trials) to enable this analysis. Of these, 16 and 19 were classified into visual and visual-movement cells, and 19 and 16 possessed and did not possess delay-period activity, respectively. In both cases, the discrimination times were significantly earlier in the Bright-Difficult than in the Dim-Easy condition (Wilcoxon signed-rank test, P < 0.01) and were not significantly different among neuron types (Wilcoxon rank-sum test, P > 0.05). This indicates that the present results were preserved regardless of neuron type.

**Behavioral and Neuronal Performances in False-alarm Error Trials**

To test whether the present findings were preserved in the data from the erroneous trials, we examined the behavioral and neuronal performances when the monkeys made false-alarm errors (fixation breaks) in the catch condition [the dataset from the erroneous trials in the search condition were not large enough for analysis due to the high rate of correct responses (e.g., >95% in the Dim-Easy search)]. We analyzed the data from the 33 neurons in which the monkeys made ≥7 false-alarm responses (erroneous saccades) to both the receptive field and the position opposite to the receptive field in each of the Bright and Dim catch conditions. The mean false-alarm error rate (± SD) was significantly higher in the Bright catch condition (55.9 ± 9.2%) than in the Dim catch condition (43.8 ± 11.5%) (signed-rank test, P < 0.0001), and the mean reaction time (± SD) of the false-alarm responses was significantly shorter in the Bright catch condition (292 ± 57 ms) than in the Dim catch condition (311 ± 62 ms) (signed-rank test, P < 0.0001), indicating that the erroneous saccades occurred more frequently and quickly in the Bright condition compared with the Dim condition.

Figure 9A shows the average population activity of the above 33 cells in the false-alarm error trials in the Bright and Dim catch conditions. The activity was greater when the false-alarm saccade goal was in the receptive field (solid traces) than when it was opposite to the receptive field (dashed traces). In the saccade-aligned activity (Fig. 9A, right), the postdiscrimination interval [the length from goal-discrimination time (signed-rank test, P < 0.01) to saccade onset] was longer in the Dim catch condition (110 ms) than in the Bright catch condition (86 ms), which is consistent with the results obtained from the successful search trials. Figure 9B shows the accumulated responses produced by applying a leaky integration function (time constant = 128 ms) to the activity of the 33 neurons. The time constant was determined in the same way as in Fig. 8B. As illustrated in the figure, the accumulated responses in both the false-alarm trials (thick lines) and the successful search trials (thin lines) converged into a common activation level at the time of saccade onset, irrespective of the stimulus condition (one-way repeated-measures ANOVA, P > 0.1), suggesting that the saccade-initiation timing on error trials is also explained by the same accumulation-to-threshold model.

**DISCUSSION**

The results from the present study reveal that the length of the postdiscrimination interval during a visual search was different when the stimulus condition was altered, even when the saccadic reaction times were matched between stimulus conditions in a trial-based manner (Figs. 3–5). Furthermore, a control analysis suggests that the postdiscrimination interval was altered by the stimulus luminance (Fig. 6). In addition, the simulation results using an accumulation-to-threshold model reveal that the across-stimulus variation in the postdiscrimination interval could be explained by the luminance-dependent changes in the strength of the response at the target-discrimination time (Fig. 8).

**Visual Stimulus Parameters Affect Pre-/Postdiscrimination Intervals in Different Ways**

As shown in Fig. 6, the two stimulus parameters (luminance and target-distractor color similarity) affected the pre- and postdiscrimination intervals in different ways. First, the pre-
discrimination interval was affected by both luminance and target-distractor similarity (Fig. 6A), but the neuronal substrate affected by these two variables might differ. Previous studies using a visual detection paradigm (in which only an isolated target was presented) revealed that luminance-level increases evoked earlier response onset and stronger visual responses in the LIP (Tanaka et al. 2013) and SC (Marino et al. 2012) and shortened the behavioral reaction time (e.g., Boch et al. 1984; Jaskowski and Sobierska 2004; Marino and Munoz 2009). In contrast, studies using a visual search paradigm (in which a target is embedded in multiple distractors) demonstrated that decreases in the target-distractor similarity did not change the visual responses but shortened the target-discrimination time and the saccadic reaction time (Bichot et al. 2001; Sato et al. 2001; Thompson et al. 2005). Therefore, magnification of the stimulus luminance might shorten the target-discrimination time by enhancing the neuronal target-distractor discriminability.

Second, the postdiscrimination interval was altered by the stimulus luminance (Fig. 6B). A similar dependence on luminance was observed in a visual detection task in which the length of time from the onset of the neuronal response to the saccade onset increased with decreasing luminance (Fig. 8 in Tanaka et al. 2013). Importantly, neurons are capable of specifying the target location at the time of response onset in a visual detection task (because only an isolated target was presented); therefore, the response-onset time in a visual detection task might correspond functionally to the target-discrimination time in a visual search task. As such, the postdetection/postdiscrimination intervals were lengthened with decreasing luminance in both the visual detection and search tasks, suggesting the existence of a common luminance-dependent process in the downstream structures.

The fact that the pre- and postdiscrimination intervals could be differentially altered under varying stimulus conditions suggests that multiple neuronal discrimination times are possible for a single behavioral reaction time. Specifically, the equivalence of behavioral reaction times does not always assume equivalent neuronal discrimination times. This suggests that, although behavioral reaction times are measured conventionally to evaluate the difficulty of a task in psychophysiological studies of perceptual discrimination, differences in behavioral reaction times do not always indicate the same differences in neuronal discrimination times in the brain under different stimulus conditions.

Relationship with Previous Studies of Saccade Target Selection in Different Visual Conditions

Although previous studies have not explicitly examined this issue, extant results indicate that the postdiscrimination interval represented in the activity of neurons in the FEF (Cohen et al. 2009) and the LIP (Balan et al. 2008) is prolonged as the number of array elements increases. Importantly, the strength of visually responsive activity decreased when the number of array elements increased, which might be caused by the effects of surround suppression (Falkner et al. 2010; Schall et al. 2004). Namely, an increase in the number of array elements would reduce the magnitude of the visual responses at the target-discrimination time. Therefore, the prolonged postdiscrimination interval observed in previous studies might be explained by the same mechanism postulated in the current study.

A previous study examined the effects of a change in luminance on SC neurons during a color-singleton search task (White and Munoz 2011). When the achromatic contrast was removed by making an array stimulus equiluminant with the background, the peak response of target activity was unremoved by making an array stimulus equiluminant with the background, whereas distractor activity was greatly attenuated. These neuronal modulations induced no change in the neuronal target-discrimination time but substantially delayed saccadic initiation to prolong the postdiscrimination interval. Therefore, prolongation of the postdiscrimination interval was unlikely to be explained by reduced response strength. This discrepancy might be due to functional differences in LIP and SC neurons or to differences in the stimulus condition; the background luminance was relatively high (20.5 cd/m²), and the array
stimulus was darker than the background in the SC study. Further studies are needed to adequately address this question.

Saccade Target Selection in Different Brain Areas

Previous studies have revealed that the profiles of target-discriminating activity during visual search vary across brain areas. First, the degree of dependence on stimulus features varies among areas. When monkeys searched for a singleton target that was unique in the instructed stimulus dimension (shape or color), target discrimination for most V4 neurons (Ogawa and Komatsu 2004) and for one-third of LIP neurons (Ogawa and Komatsu 2009) occurred only when the target was defined by one particular dimension. However, half of the LIP neurons and most FEF neurons discriminated the target, irrespective of the target-defining dimension (Ogawa and Komatsu 2006, 2009). Consistently, the percentage of shape-selective neurons was significantly lower in the FEF compared with the LIP (Lehky and Sereno 2007; Peng et al. 2008). Second, saccadic reaction times covaried with the target-discrimination time for neurons in the LIP and FEF (Bichot et al. 2001; Ipatu et al. 2006; Sato et al. 2001; Thomas and Paré 2007; but see Thompson et al. 1996) and for some neurons in the SC (McPeek and Keller 2002) but not for V4 neurons (Gee et al. 2010). These differences in the target-discriminating response profiles may reflect the different hierarchies in the transition from stimulus-dependent to spatially based target discrimination (Gottlieb and Snyder 2010). Taken together with the present finding that the postdiscrimination interval can differ across stimulus conditions, it is suggested that the temporal relationship between the target-discriminatory interval and the saccade-initiation time may vary not only across stimulus conditions but also among brain areas.

Discrete vs. Continuous Flow Models

Two conceptual models have been suggested to explain the nature of sensorimotor transformation: discrete- and continuous-flow models (Costello et al. 2013; Eriksen and Schultz 1979; Miller 1982; Smid et al. 1991; Woodman et al. 2008). A discrete-flow model assumes that different independent stages are processed in a sequential manner, which is consistent with the conception of a subtraction method (Donders 1868/1969). This model proposes that target discrimination and saccade preparation/generation are independently and discontinuously processed, and it therefore predicts that the postdiscrimination interval will be the same regardless of stimulus condition. In contrast, a continuous-flow model allows that the effects of the stimulus remain even after the target-discriminatory time, and it therefore predicts that the neuronal activity during the postdiscrimination interval will differ across different stimulus conditions. Thus, the current results are more likely explained by a continuous-flow model rather than by a discrete-flow model.

Woodman et al. (2008) examined the same problem by recording the activity of FEF movement neurons during a visual search and proposed the opposite conclusion. Nevertheless, we think that their and our conclusions can be reconciled. As proposed recently (Purcell et al. 2012), if the inputs from visual neurons to movement neurons are gated until the target discriminability achieves a threshold, movement neurons do not start the activation. In such a situation, LIP visual neurons behave in a continuous-flow manner, whereas FEF movement neurons behave in a discrete-flow manner. Thus, it is possible that continuous- and discrete-flow processing occur concurrently in the activity of the different types of neuron in the brain.

Exogenous and Endogenous Influences on the Postdiscrimination Interval

The results from the present study suggest that visual stimulus (luminance) affected the determination of the postdiscrimination interval. In addition to this exogenous factor, previous studies have suggested that endogenous factors also affect the postdiscrimination interval. Although it is known that the length of time from target discrimination to saccadic onset tends to be constant despite the variability of the saccadic reaction time in a given stimulus condition (Ipatu et al. 2006; Sato et al. 2001; Thomas and Paré 2007), the degree of this constancy could be modified by differences in the task design of the visual search. When monkeys were required to foveate the target by a single saccade in the visual search (slow-search-speed/high-search-accuracy condition), constancy in the postdiscrimination interval was observed in approximately half of the visual movement neurons in the SC (McPeek and Keller 2002). In contrast, when monkeys were allowed to foveate the target by multiple saccades (fast-search-speed/low-search-accuracy condition), constancy in the postdiscrimination interval was observed across the entire population of SC neurons (Shen and Paré 2011). In addition, a recent study reported that when a speed-accuracy tradeoff was manipulated using a prestimulus instruction cue, the length of the postdiscrimination interval increased with search accuracy even when an identical array stimulus was presented (Heitz and Schall 2012). These results suggest that the length of the postdiscrimination interval could be altered by endogenous factors.

In the current study, the reaction-time trials were randomly interleaved with no explicit instruction, and the monkeys were unable to anticipate the upcoming stimulus condition. Furthermore, the present results demonstrate that the length of the postdiscrimination interval differed even when the saccadic reaction times were almost perfectly matched between the two different stimulus conditions, suggesting that the present results could not be explained by the above endogenous factors. Nevertheless, it is still possible that the monkeys rapidly altered their search speed/search accuracy within a trial after seeing a presented array stimulus. The monkeys might strengthen their speed-stress response after seeing an Easy stimulus compared with a Difficult stimulus, because they would have had a higher probability of getting a reward due to the increased correct response rate in the Easy compared with the Difficult condition (Fig. 1). However, this was not the case because the postdiscrimination interval was not statistically different between the Easy and Difficult conditions at each luminance level (Fig. 6B). Taken together, the present findings suggest that changes in the postdiscrimination interval were likely not due to the endogenous factors but, rather, were due to exogenous factors (the visual stimulus).

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DISCLOSURES

No conflicts of interest, financial or otherwise, are declared by the author(s).

AUTHOR CONTRIBUTIONS


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