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Quantification of Fractional Dynamical Stability of EEG Signals as a Bio-Marker for Cognitive Motor Control

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Assessing the stability of biological system models has aided in uncovering a plethora of new insights in genetics, neuroscience, and medicine. In this paper, we focus on analyzing the stability of neurological signals, including electroencephalogram (EEG) signals. Interestingly, spatiotemporal discrete-time linear fractional-order systems (DTLFOS) have been shown to accurately and efficiently represent a variety of neurological and physiological signals. Here, we leverage the conditions for stability of DTLFOS to assess a real-world EEG data set. By analyzing the stability of EEG signals during movement and rest tasks, we provide evidence of the usefulness of the quantification of stability as a bio-marker for cognitive motor control.

Keywords: nonlinear models, stability, EEG signals, control applications, cognitive motor control

1 INTRODUCTION

Systems biology and biomedicine has been studied in conjunction with control theory over several decades (Wiener, 2019). By applying the tools developed in control theory to biological and biomedical systems, new discoveries have been made that enhance our understanding of both the function and dysfunction of biological systems. One such control theoretic tool is stability, which has been particularly useful in uncovering new critical insights in a multitude of biological and biomedical applications (Wang and Zhang, 2009; Zhang et al., 2015; Li et al., 2017). Stability assesses the long-term behavior of a dynamical model and its equilibrium properties, which are critical to understanding the nature of the underlying biological or biomedical process and its characteristics. However, the accuracy of such conclusions drawn on the properties of the biological or biomedical process under observation will depend on the quality-of-fit of the model that is used to describe the underlying process. Assessing the goodness-of-fit requires measurements of the biological process.

There are many different tools to measure neurophysiological signals, including electrocardiogram (ECG), electromyogram (EMG), electroencephalogram (EEG), and blood oxygenation level dependent (BOLD), and their models have been studied in the literature (Baleanu et al., 2011; Magin, 2012; Pequito et al., 2015; Xue et al., 2016b; Gupta et al., 2018). In this paper, we restrict our attention to non-invasive scalp EEG, which has been widely used to draw conclusions on disease diagnosis, develop treatment, and gain a fundamental understanding of systems biology on the whole (Haddad et al., 2010; McFarland et al., 2010).

There is evidence that *discrete-time linear fractional-order dynamical systems* (DTLFOS) are well-suited to describe EEG signals and forecast their evolution (Pequito et al., 2015; Gupta et al., 2018). Furthermore, these nonlinear *spatiotemporal* models rely on only a few parameters

(Caponetto, 2010; Xue et al., 2016b; Monje et al., 2010). These parameters include a *coupling* matrix and a set of temporal scale coefficients called *fractional exponents*. The coupling matrix describes the spatial dependencies between different state variables. The fractional exponents describe the long-term memory of the dynamical process across different state variables. With these few required and easy-to-interpret parameters, together with the superior goodness-of-fit, these DTLFOS models present distinct advantages over both linear time-invariant and (generically) nonlinear models. Specifically, LTI systems are limited in their ability to capture different time-scales occurring in biological applications, which are inherently modeled by the spatial component (i.e., the system's autonomous matrix) (Pequito et al., 2015; Xue et al., 2016a; Xue and Bogdan, 2017). On the other hand, spatiotemporal nonlinear systems have been widely accepted as tools capable of accurately modeling EEG data but may lead to over-complicated and difficult to interpret models that often over-fit the data.

Despite the advantages of DTLFOS, there are few results regarding the stability of fractional-order systems (FOS). The prior literature describes conditions for continuous-time FOS (Benzaouia et al., 2014) and for single-input single-output continuous-time commensurate systems (i.e., systems with equal fractional exponents across state variables) (Dastjerdi et al., 2019). In Dzieliński and Sierociuk (2008), the authors leverage an infinite dimensional representation of truncated DTLFOS (i.e., with finite memory) to give conservative sufficient conditions for stability. While the work in (Busłowicz and Ruszewski, 2013) does provide necessary and sufficient conditions for practical and asymptotic stability of DTLFOS, they only consider commensurate-order systems (i.e., α is the same for all state variables). That said, a comprehensive analysis of DTLFOS stability is lacking.

Henceforth, we aim to analyze the quantification of stability of DTLFOS that encode EEG signals giving us the ability to unveil new insights on the relation between neural behavior and cognitive motor control. The prior literature describes methods using EEG signals to classify various movements with varying levels of success (Morash et al., 2008; Kim et al., 2019; Ofner et al., 2019). However, to the best of our knowledge, analyzing the stability of DTLFOS that model EEG signals to draw conclusions on cognitive motor control is a novel approach.

Subsequently, the main contributions of this work are as follows. First, we introduce stability conditions for multivariate DTLFOS with arbitrary fractional exponents. We then leverage these conditions to study the stability of a real-world EEG motor data set modeled as a DTLFOS and provide evidence of its relevance in the context of cognitive motor control.

2 MATERIALS AND METHODS

2.1 Discrete-Time Linear Fractional-Order System

In this paper, we consider discrete-time linear fractional-order non-commensurate systems described by

$$\Delta^\alpha x[k+1] = Ax[k], \quad (1)$$

where $x \in \mathbb{R}^n$ denotes the state and Δ^α is the Grünwald-Letnikov discretization of the fractional derivative. The Grünwald-Letnikov discretization for any i -th state ($1 \leq i \leq n$) can be expressed as $\Delta^\alpha x_i[k] = \sum_{j=0}^k \psi(\alpha_i, j)x_i[k-j]$, where $\alpha_i \in \mathbb{R}$ is the fractional-order coefficient of the i th state and $\psi(\alpha_i, j) = \frac{\Gamma(j-\alpha_i)}{\Gamma(-\alpha_i)\Gamma(j+1)}$ with $\Gamma(\cdot)$ denoting the Gamma function (Dzieliński and Sierociuk, 2005).

Notice that System (Eq. 1) is a (finite-dimensional) nonlinear (non-Markovian) system, and it considers a weighted linear combination of all the previous data from the beginning to the current time. As can be noticed from the Grünwald-Letnikov discretization formula, the magnitude of the fractional-order derivative necessitates the total number of previous data points and the weights on those data points. Intuitively, a larger fractional-order coefficient (i.e., α closer to 1) implies a lower dependency on the previous data meaning that the weights decay at a faster rate. Furthermore, as the fractional-order coefficient α approaches 1, the system becomes closer to an integer-order linear time-invariant system.

Towards deriving the stability conditions for DTLFOS, we start by noticing that the DTLFOS (Eq. 1) can be re-written as [Lemma 2, (Gupta et al., 2018)]:

$$x[k] = G_k x[0], \quad (2)$$

where

$$G_k = \begin{cases} I_n, & k = 0 \\ \sum_{j=0}^{k-1} A_j G_{k-1-j}, & k \geq 1 \end{cases} \quad (3)$$

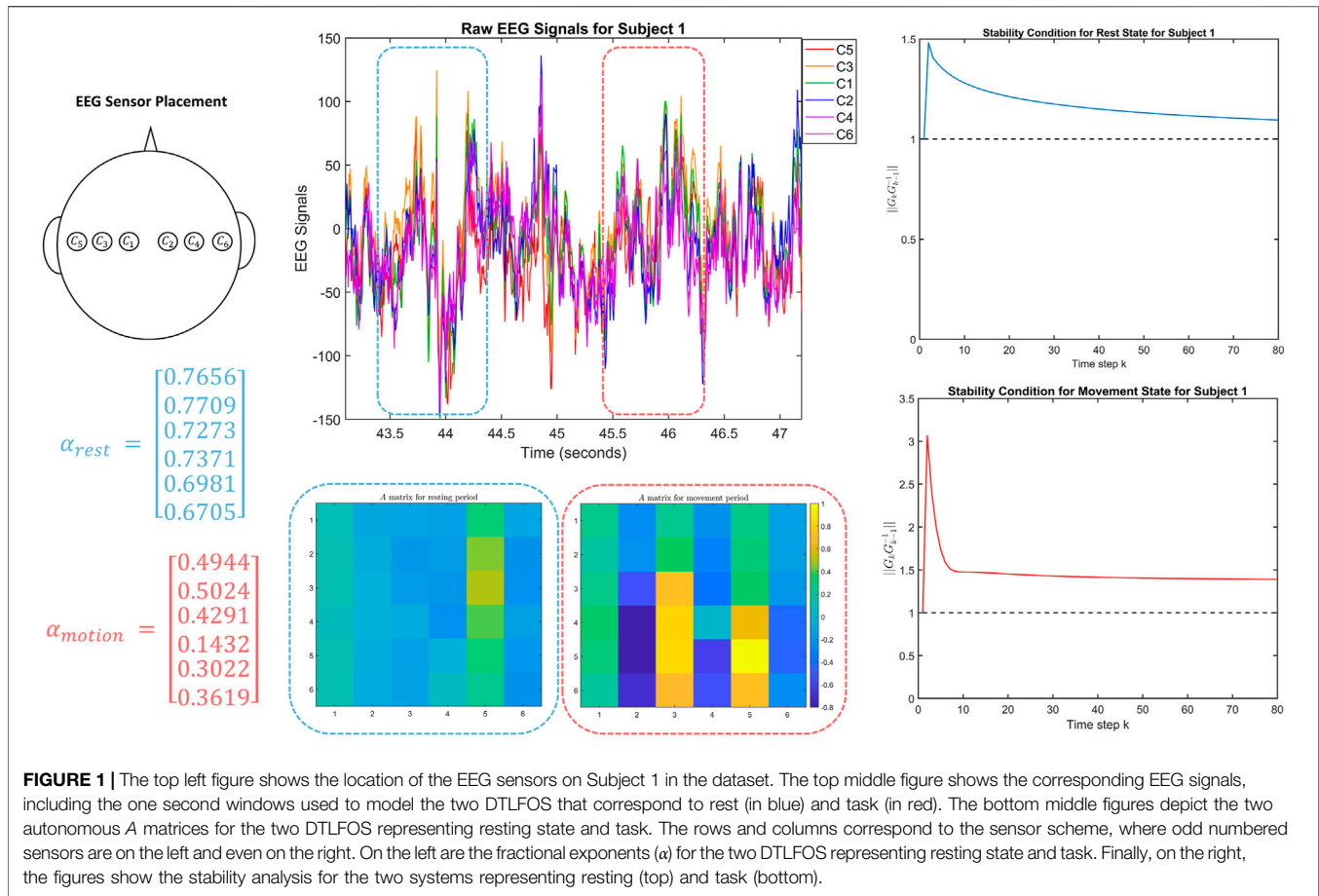
with $A_0 = A - D(\alpha, 1)$, $A_j = -D(\alpha, j+1)$, for $j \geq 1$, and

$$D(\alpha, j) = \begin{bmatrix} \psi(\alpha_1, j) & 0 & \dots & 0 \\ 0 & \psi(\alpha_2, j) & \dots & 0 \\ 0 & \vdots & \ddots & 0 \\ 0 & 0 & \dots & \psi(\alpha_n, j) \end{bmatrix}. \quad (4)$$

In particular, we observe that Eq. 2 will lead to the following realizations.

$$\begin{aligned} G_0 &= I_n \\ G_1 &= A - D(\alpha, 1) \\ G_2 &= A^2 - AD(\alpha, 1) - D(\alpha, 1)A + D(\alpha, 1)^2 - D(\alpha, 2) \\ G_3 &= A^3 - AD(\alpha, 1) - D(\alpha, 1)A^2 + D(\alpha, 1)^2 \\ &\quad - A^2D(\alpha, 1) + AD(\alpha, 1)^2 + D(\alpha, 1)AD(\alpha, 1) \\ &\quad + D(\alpha, 1)^3 - AD(\alpha, 2) + D(\alpha, 1)D(\alpha, 2) \\ &\quad - D(\alpha, 2)A + D(\alpha, 2)D(\alpha, 1) - D(\alpha, 3) \\ &\vdots \end{aligned} \quad (5)$$

Using Equation 2, we show that a DTLFOS as in Eq. 1 can be represented as a discrete-time linear time-varying (DTLTV) system under mild assumptions. First, we notice that a DTLTV is described by



$$x[k+1] = M_k x[k], \quad (6)$$

with a state-transition matrix (that follows from using Eq. 6 recursively)

$$x[k] = \left(\prod_{i=0}^{k-1} M_{k-1-i} \right) x[0], \quad (7)$$

which provides a mapping from the initial state to the state at any time k . Consequently, from Eqs 2, 7, we find the conditions for which a DTLFOS can be represented as a DTLTV by equating the state-transition matrices $G_k = \prod_{i=0}^{k-1} M_{k-1-i}$, $\forall k$.

Next, we observe that $M_k = G_{k+1} G_k^{-1}$ under the assumption that G_k is invertible for all k . In particular, we notice that $\prod_{i=0}^{k-1} M_{k-1-i} = \prod_{i=0}^{k-1} G_{k-i} G_{k-1-i}^{-1} = (G_k G_{k-1}^{-1})(G_{k-1} G_{k-2}^{-1})(G_{k-2} G_{k-3}^{-1}) \dots G_1 G_0^{-1} = G_k G_0^{-1} = G_k$.

Notice that the assumption on the invertibility of G_k for all k can be seen as a mild assumption by invoking measure theoretical arguments. Simply speaking, consider any random matrix A , which is invertible almost surely since having a non-invertible A would require the determinant of A to be zero, which has a zero probability of occurring. From Eq. 5, we see that there would need to be a very specific combination to make G_k have a determinant of zero and thus be non-invertible, which again occurs with probability zero. Since the specific combinations required to make G_k have a zero

determinant are different for each k , we do not define the specific combination; however, we can give some intuition as to what this combination might be by providing some examples. We remark that for G_1 , the diagonal entries of $D(\alpha, 1)$ would need to be such that when subtracted from A , the matrix $A - D(\alpha, 1)$ is not invertible. Similarly for G_3 , we see that there needs to be specific combinations of the matrices A , $D(\alpha, 1)$, $D(\alpha, 2)$, $D(\alpha, 3)$ and any certain powers of these to attain a non-invertible matrix G_3 . From these examples, we see that the specific combination for a matrix G_k to be non-invertible is dependent on A , $D(\alpha, i)$, and any powers of these up to i , where $i = \{1, \dots, k\}$.

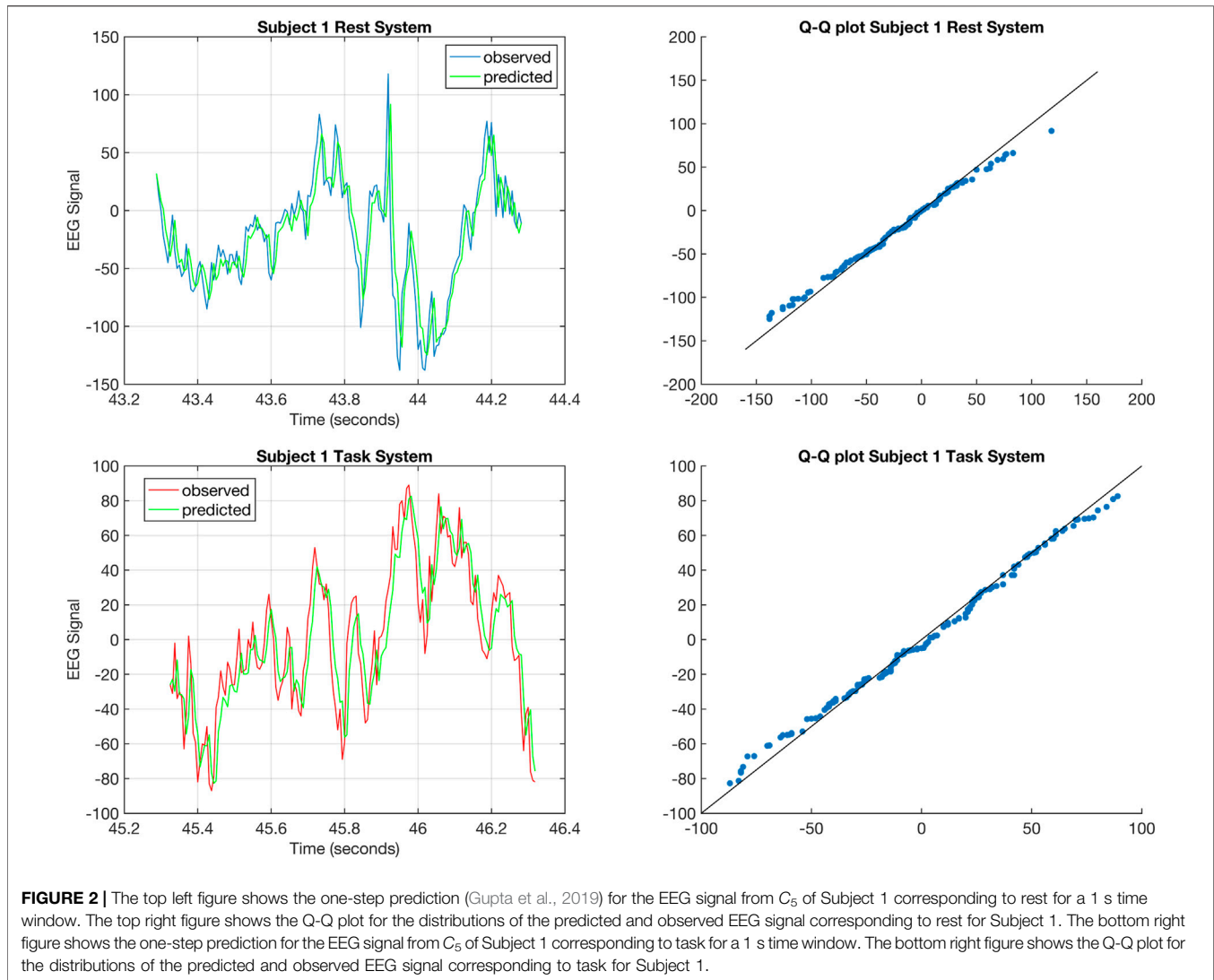
By showing that the DTLFOS can be re-cast as a DTLTV system under certain assumptions, we can now leverage the stability results from DTLTV systems to derive a stability condition for DTLFOS.

We start by introducing the definition of stability.

Theoretical Stability: (Section 4.1, (Khalil, 2014)) A DTLFOS system (Eq. 1) where G_k is invertible for all k system is said to be

- stable if for any $\epsilon > 0$ and for any k_0 , there exists a $\delta(k_0, \epsilon) > 0$ such that $\|x[k_0]\|_2 \leq \delta(k_0, \epsilon) \Rightarrow \|x[k]\|_2 \leq \epsilon, \forall k \geq k_0$.
- unstable if it is not stable.

In the main result, we provide the necessary and sufficient conditions of stability for DTLFOS (1).



Theorem 1. A DTLFOS (1), where G_k is invertible for all $k \in \mathbb{N}$, is stable if and only if there exists some constant $0 < c(k_0) < \infty$ such that

$$\|G_k G_{k-1}^{-1}\|_2 \leq c(k_0), \forall k, k_0, \quad (8)$$

where $k \geq k_0$.

PROOF. Assuming that G_k is invertible for all $k \in \mathbb{N}$, it follows that we can represent the DTLFOS as a LTV system, i.e., $x[k+1] = M_k x[k]$, where $M_k = G_{k+1} G_k^{-1}$. Leveraging the fact that the DTLFOS can be represented as a LTV, we know that a discrete-time LTV system is stable if and only if for any k_0 , there exists a $0 < \gamma(k_0) < \infty$, such that $\|\Phi(k, k_0)\|_2 \leq \gamma(k_0)$, $\forall k, k_0$, where $k \geq k_0$ and $\Phi(k, k_0)$ is the state-transition matrix from the state at k_0 to the state at k [Lemma 1, (Zhou and Zhao, 2017)]. Therefore, to show sufficiency, we notice that

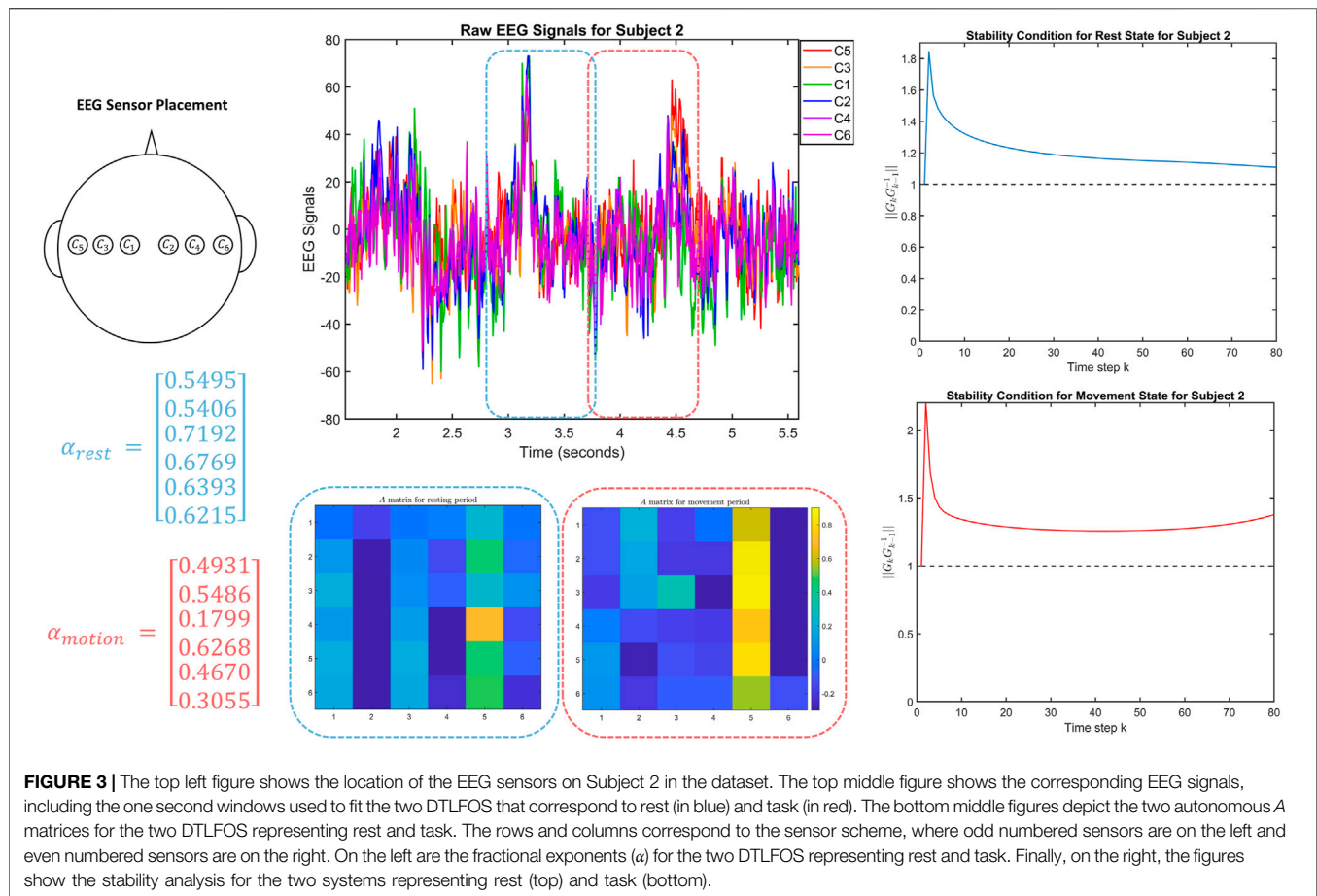
$$\begin{aligned} \|\Phi(k, k_0)\|_2 &= \|M_{k-1} \dots M_{k_0}\|_2 \\ &\leq \|G_k G_{k-1}^{-1}\|_2 \|G_{k-1} G_{k-2}^{-1}\|_2 \dots \|G_{k_0+1} G_{k_0}^{-1}\|_2 \\ &\leq c(k_0)^{k-k_0} = \gamma(k_0). \end{aligned}$$

The necessary condition can be shown by contradiction. Suppose that there exists an index k_i for which there does not exist a $0 < c(k_0) < \infty$ such that $\|G_{k_i} G_{k_i-1}^{-1}\|_2 \leq c(k_0)$, then it must mean that $\|G_{k_i} G_{k_i-1}^{-1}\|_2 > \infty$, so $\|\Phi(k_i, k_0)\|_2 > \infty$, so the system is unstable.

We define the quantification of stability at time k in the following definition.

Definition 1. The quantification of stability at time k is defined by the magnitude of the stability metric $\|G_k G_{k-1}^{-1}\|_2$ given that $\|G_k G_{k-1}^{-1}\|_2 < \infty$.

Given EEG signals during a period when a subject is resting and when s/he is performing task, we will fit these signals to two distinct DTLFOS for the rest and task periods by using the method outline in Xue et al. (2016b), which finds the parameters using least-squares regression. By comparing the stability and quantifying the stability metric $\|G_k G_{k-1}^{-1}\|_2$ of these two systems, we expect to observe differences that not only help distinguish these two activities but also aid in



establishing and verifying new and existing conclusions in cognitive motor control.

3 ASSESSING THE QUANTIFICATION OF STABILITY OF EEG SIGNALS AS A BIO-MARKER FOR COGNITIVE MOTOR CONTROL

In this section, we leverage the stability of DTLFOS systems (Eq. 1) to assess EEG signals for motor movement. DTLFOS systems are stable if and only if $\|G_k G_{k-1}^{-1}\|_2$ is bounded by a finite constant for all time steps k .

We explore the stability of EEG signals in the motor cortex for 106 subjects completing simple motor tasks. In each trial, a target

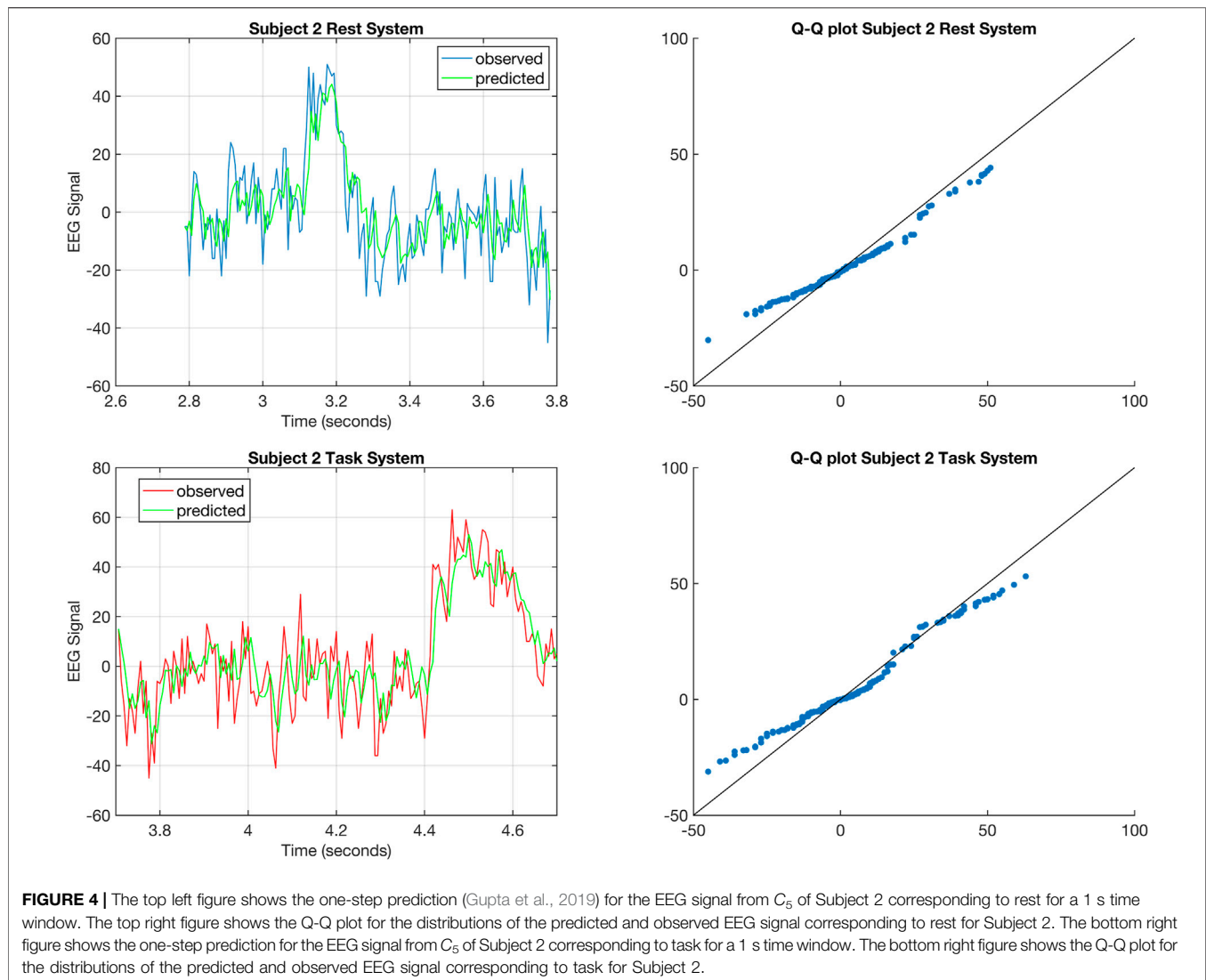
appears on the left side of a screen, at which point the subject opens and closes the left fist until the target disappears, then the subject relaxes (Schalk et al., 2004). This data set consists of over 1,500 trials corresponding to one- and 2-min EEG recordings, obtained from 106 volunteers (Goldberger et al., 2000).

We use the data collected from the six sensors over-top the motor cortex to fit two distinct DTLFOS (Xue et al., 2016b), one corresponding to when the subject is resting and the other corresponding to when the subject is completing the motor task. For each subject, we chose a one second window of time at which point the subject is resting and a subsequent one second window when the subject begins to perform the task (open and close the left fist). Using these two windows of data, we fit two DTLFOS, one corresponding to the resting period and the other corresponding to the movement task. With these two systems, we use **Algorithm 1** to evaluate the stability of the DTLFOS for 160 time-steps. We note that if the stability metric, namely $\|G_k G_{k-1}^{-1}\|_2$ exceeded a threshold of 300, we replaced it with the value 300. We note that for the selected time windows all of the values of $\|G_k G_{k-1}^{-1}\|_2$ were less than infinity, so all systems were found to be stable. The time window should be selected carefully to avoid numerical instability (Xue et al., 2016b).

Briefly, we note that in general, these systems are stable. The stability metric $\|G_k G_{k-1}^{-1}\|$ captures the overall activity of the motor cortex. More specifically, $\|G_k G_{k-1}^{-1}\|$ over time captures

Algorithm 1 | This algorithm is used to assess the stability of the DTLFOS (1) for a finite time horizon.

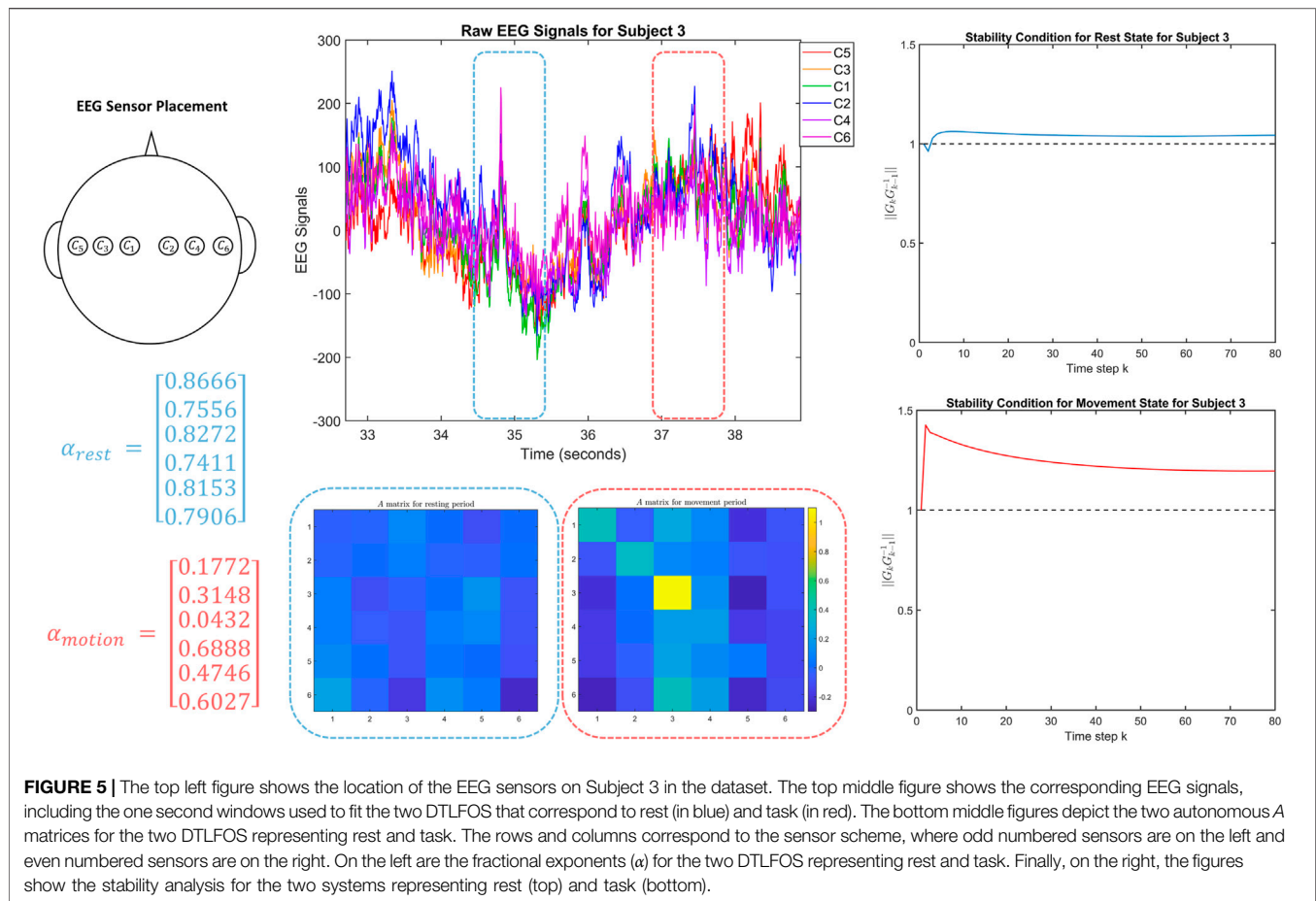
Input: Given K, α, A
Initialization: Set $k = 1$, compute the state-transition matrix G_k for $k = 1, \dots, K$ using (3);
while $k < K$ **do**
 if $\|G_k G_{k-1}^{-1}\|_2 > \infty$ **then**
 Output: the DTLFOS (1) is unstable;
 break;
 end if
 end while
Output: the DTLFOS (1) is stable up to finite time K ;



the qualitative change of behavior of the underlying system as seen in the transient of the stability metric for Subjects 1, 2, and 3. We notice that during the task, the system has larger $\|G_k G_{k-1}^{-1}\|_2$ values when compared to the resting system. Previous work has shown that the neural activity in the motor cortex increases as muscles contract to perform a task (Chapter 37, Kandel et al. (2000)). This might explain why the task DTLFOS has higher $\|G_k G_{k-1}^{-1}\|_2$ values when compared to the resting DTLFOS since the higher brain activity while performing a task might be associated with an increase in $\|G_k G_{k-1}^{-1}\|_2$. Therefore, by observing that the stability metric quantitatively changed when comparing the rest system to the task system, this suggests that the behavior of the underlying system also changes. The larger $\|G_k G_{k-1}^{-1}\|_2$ values for the task system indicate that the quantification of the stability of DTLFOS can serve as a *bio-marker* for the brain's change in behavior when switching from resting to movement tasks. We provide evidence to support this claim both by exploring several subjects and by statistically

analyzing all 106 subjects through a Kolmogorov-Smirnov test (Hogg et al., 2010). The results for Subjects 1, 2, and 3 are presented in **Figures 1, 3, 5**, respectively. Furthermore, we evaluate the quality of fit by performing a one-step prediction for the rest and task DTLFOS and their corresponding Q-Q plots for Subjects 1, 2, and 3 in **Figures 2, 4, 6**, respectively.

In **Figure 1**, we observe several yellow squares appearing in the autonomous A matrix during the task. This is synonymous with the right side of the brain being activated during the movement of the left fist, which matches the conclusions of movement drawn by neuroscientists. These conclusions state that the right side of the brain controls the left side of the body and vice versa. Also during the task, we notice that the fractional exponents (α) are closer to 0.5 indicating that the system depends more heavily on information from signals in the very distant past. Therefore, a higher weight is being placed on previous states in the task system. However, in the resting system, the fractional exponents are closer to 1 indicating that the system does not



depend on signals in the very distant past and requires less dependence on the previous states. The fact that the rest system does not depend heavily on previous states may be due to a possible transition between different cognitive states as expected in default mode (Breakspear, 2017).

In **Figure 2**, we evaluate the quality of fit by performing a one-step prediction for the rest and task DTLFOS and their corresponding Q-Q plots for Subject 1. The Q-Q plots shows that the fit is good when the points exhibit a linear relationship (Hogg et al., 2010). **Figure 2** provides evidence that the estimated DTLFOS is a good fit for the EEG signals.

In the stability metric plots on the right side of **Figure 1**, the rest system and task system tend towards stability as the time-step increases, which is likely due to the fact that we are modeling an excited system driven by the inputs. As such, the behavior appearing in the stability analysis is similar to what would happen when applying an impulse or step input to a system, where we notice an initial transient followed by a return to stability.

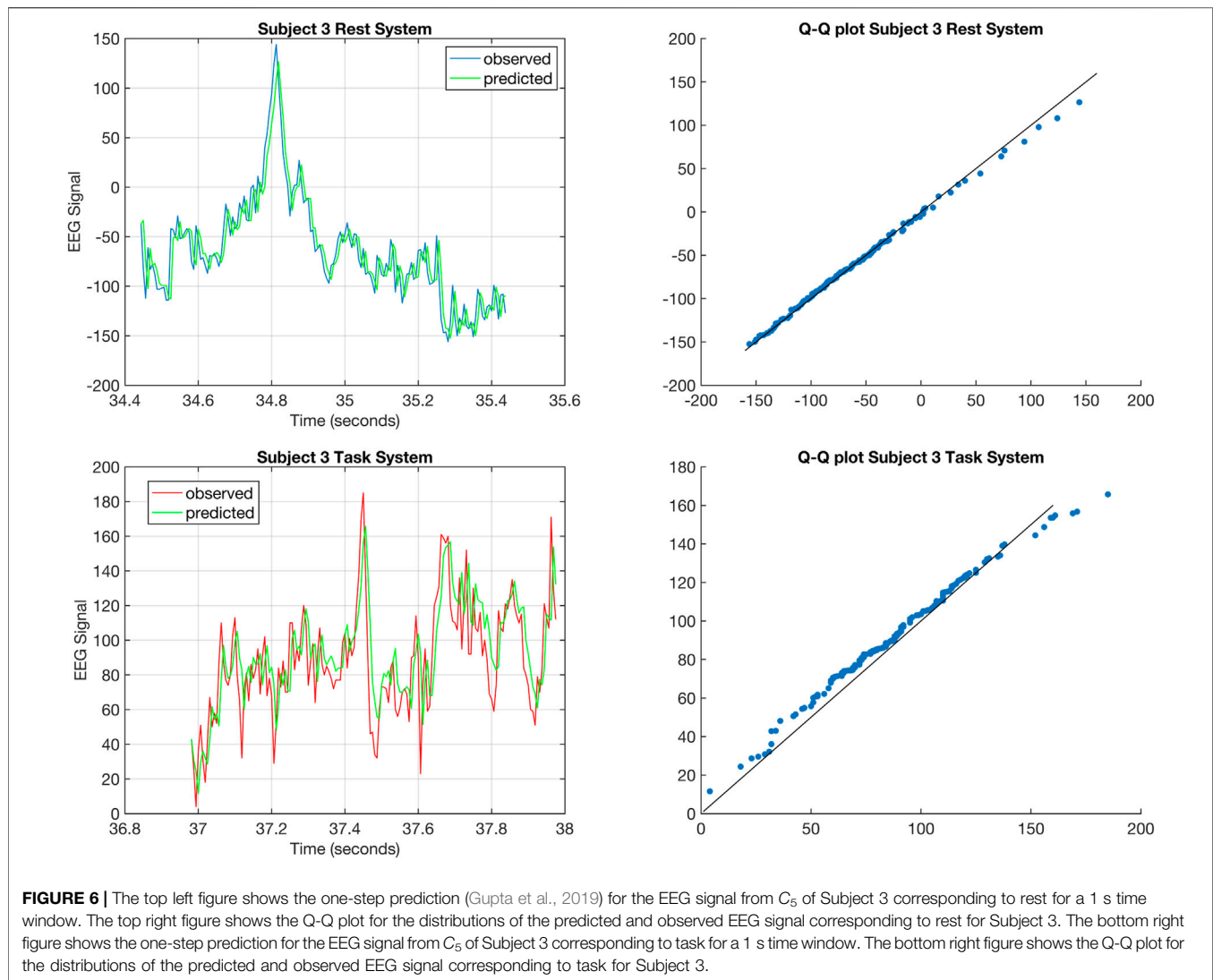
We notice in the rest system that the value of $\|G_k G_{k-1}^{-1}\|_2$ peaks around 1.5, whereas in the task system the value of $\|G_k G_{k-1}^{-1}\|_2$ peaks around 3. In general the values of $\|G_k G_{k-1}^{-1}\|_2$ are larger for the task system than the rest system. Hence, we conclude that the quantification of stability of DTLFOS can

help in providing further intuitions about the dynamics of the brain and serve as a bio-marker for its change in behavior.

In **Figure 3**, we see similar results to that of Subject 1, where during the task, the right side of the brain is activated and the fractional exponents are lower indicating that the task system depends heavily on previous states. We also see a similar behavior in the stability analysis where $\|G_k G_{k-1}^{-1}\|_2$ is larger during the task and reaches a peak around 2.2, whereas in the rest system, the peak is around 1.8.

While the Q-Q plots in **Figure 4** do not exhibit a truly linear relationship, we argue that the results from Subject 2 still produce good results and so the fit of the estimated DTLFOS to the EEG signals is still fairly good.

Finally, in **Figure 5** while despite being similar to that of Subjects 1 and 2, the results are less dramatic. However, we again note the same trends of the fractional exponents and the stability metrics for rest and task. In particular, the fractional-order exponents are lower for the task than for the rest system. Furthermore, the peak value for $\|G_k G_{k-1}^{-1}\|_2$ is around 1.4 for the task system and 1.1 for the rest system. The activation in the autonomous matrix is evident but much less pronounced and occurs in the middle of the motor cortex as opposed to on the presumed right side. Still, the results from Subject 3



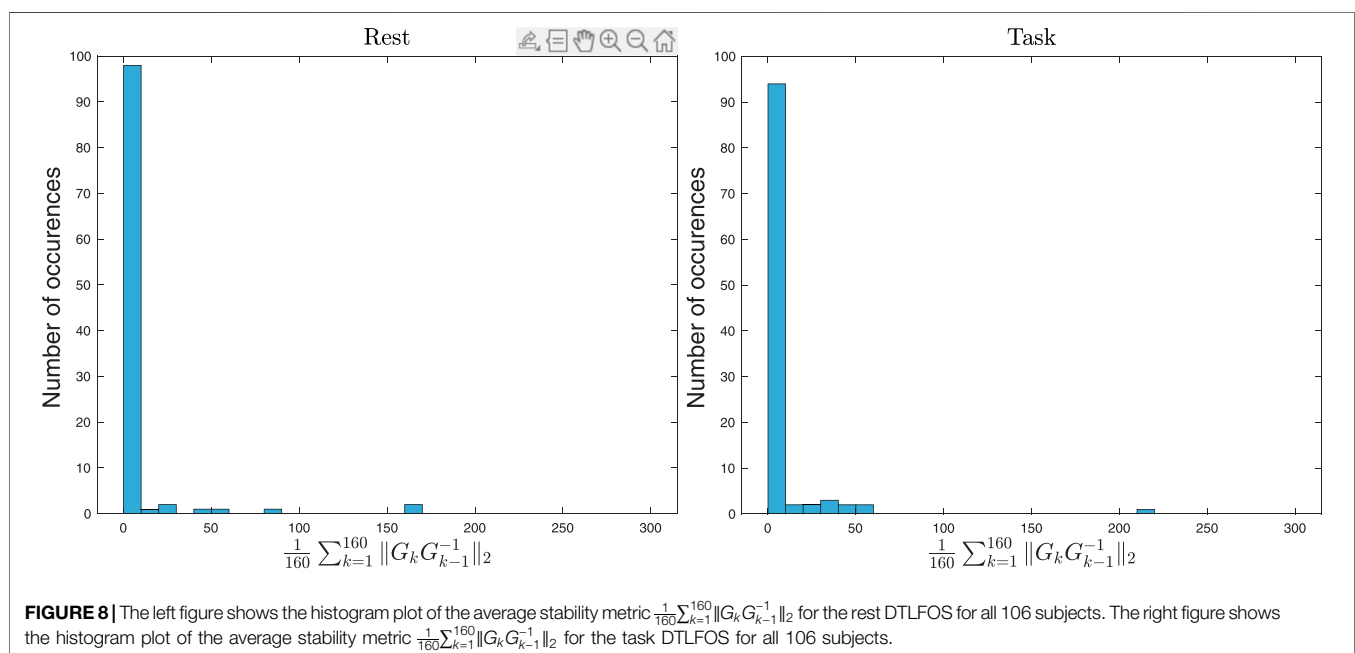
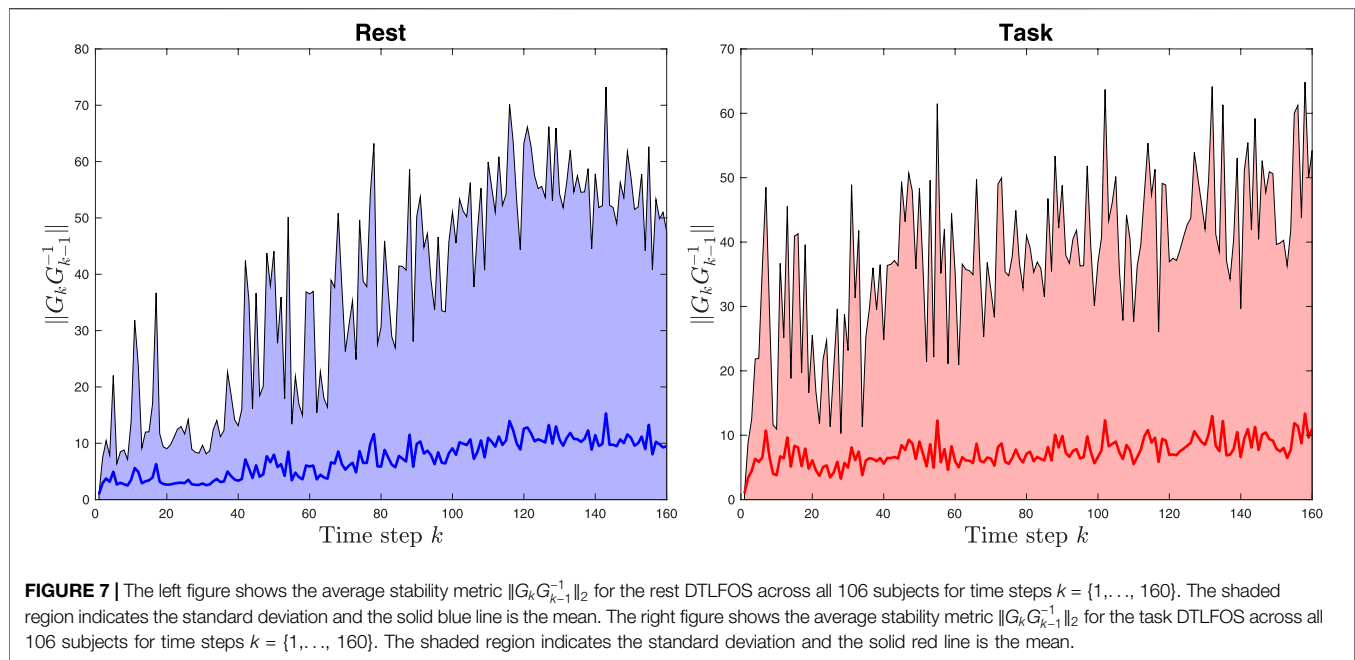
support the fact that significant and important conclusions regarding motor movement, including the transition from rest to task, can be drawn from determining the quantification of stability of the EEG signals when fit as a DTLFOS. Furthermore, the quality of fit of the DTLFOS to the EEG signals is very good for Subject 3 as evidenced by the plots shown in **Figure 6**.

Figure 7 show the average stability metric ($\|G_k G_{k-1}^{-1}\|_2$ for $k = 1 \{1, \dots, 160\}$) across all 106 subjects during rest and task periods, respectively. It is important to point out that the average stability metric is larger during the task period than the rest period. Next, we provide statistical evidence that quantifying the average stability metric of EEG signals for rest and movement tasks indicates a change in the brain's behavior and subsequently a bio-marker for cognitive motor control.

We use the Kolmogorov-Smirnov test to show that indeed the distributions of the average values of the stability metric across 160 time steps ($\frac{1}{160} \sum_{k=1}^{160} \|G_k G_{k-1}^{-1}\|_2$ for the rest and task systems are

significantly different. We start by finding the histogram plots of the average values of the stability metric across 160 time steps ($\frac{1}{160} \sum_{k=1}^{160} \|G_k G_{k-1}^{-1}\|_2$) for the rest and task systems for all 106 subjects in **Figure 8**. Next, we find the Kolmogorov-Smirnov statistic to be 0.0943. With an α level of 0.001, we find that the Kolmogorov-Smirnov statistic rejects the null hypothesis, meaning that the two distributions are statistically different. In particular, the rest distribution has more lower average values of the stability metric than the task distribution. This supports the conclusion found in Subjects 1-3 that the quantification of the stability of EEG signals modeled as a DTLFOS during rest and task are different, where the values of $\|G_k G_{k-1}^{-1}\|_2$ are generally larger for the task system when compared with the rest system.

We find that the results hold for different window sizes. When considering a window size of 0.875 s, the Kolmogorov-Smirnov statistic is 0.0849, so the null hypothesis is rejected with an α level of 0.001. Similarly, when considering a window size of 1.25 s, the Kolmogorov-Smirnov statistic is 0.0755, so again the null



hypothesis is rejected with an α level of 0.001. Hence, we provided evidence that the quantification of stability, namely how large the stability metric $\|G_k G_{k-1}^{-1}\|_2$ is compared to the baseline, can be used as a bio-marker for cognitive motor control, where the baseline is the subject at rest. This is significant because the quantification of stability can be used to design a discriminator for classifying EEG signals corresponding to rest or task.

In summary, this real-world example of fitting EEG signals to DTLFOS models and determining their quantification of stability leads to a deeper understanding of brain dynamics and motor movement.

4 CONCLUSIONS AND FUTURE WORK

The quantification of stability plays a key role in unveiling important characteristics of biological systems. Notably, spatiotemporal discrete-time linear fractional-order systems can accurately and efficiently represent a variety of biological networks. We focused on physiological systems, especially EEG signals, and we leveraged the stability conditions for DTLFOS introduced in this paper to identify features that could serve as biomarkers for cognitive motor control. Specifically, we studied a real-world EEG data set consisting of 106 subjects and fit two different DTLFOS

corresponding to the recordings during a subject's rest and movement. We showed that analyzing the quantification of stability of these two systems is useful in distinguishing a change in the brain's behavior, which can help enhance our understanding of motor movement and control.

In future work, we seek to determine necessary and sufficient conditions for stability of DTLFOS that are computationally tractable. In addition, we want to understand the trade-offs between the stability of the system and the memory captured by the system's temporal parameters (fractional-order exponents), which will ultimately lead to a better understanding of how memory affects cognitive motor control and also neurological diseases (e.g., epilepsy).

DATA AVAILABILITY STATEMENT

The original contributions presented in the study are included in the article/**Supplementary Material**, further inquiries can be directed to the corresponding author. The dataset analyzed for this study can be found in the <https://physionet.org/content/eegmmidb/1.0.0/>

AUTHOR CONTRIBUTIONS

ER. derived the theoretical conditions and completed the experiments. PB. contributed to the development of the theoretical results and experiments, and revised the paper.

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SUPPLEMENTARY MATERIAL

The Supplementary Material for this article can be found online at: <https://www.frontiersin.org/articles/10.3389/fcteg.2021.787747/full#supplementary-material>

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