

1 Real-time detection of spoken speech from unlabeled 2 ECoG signals: A pilot study with an ALS participant

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22 Abstract

23 *Objective.* Brain-Computer Interfaces (BCIs) hold significant promise for restoring communication in
24 individuals with partial or complete loss of the ability to speak due to paralysis from amyotrophic lateral
25 sclerosis (ALS), brainstem stroke, and other neurological disorders. Many of the approaches to speech
26 decoding reported in the BCI literature have required time-aligned target representations to allow
27 successful training – a major challenge when translating such approaches to people who have already lost
28 their voice. *Approach.* In this pilot study, we made a first step toward scenarios in which no ground truth

29 is available. We utilized a graph-based clustering approach to identify temporal segments of speech
30 production from electrocorticographic (ECoG) signals alone. We then used the estimated speech
31 segments to train a voice activity detection (VAD) model using only ECoG signals. We evaluated our
32 approach using held-out open-loop recordings of a single dysarthric clinical trial participant living with
33 ALS, and we compared the resulting performance to previous solutions trained with ground truth acoustic
34 voice recordings. *Main results.* Our approach achieves a median error rate of around 0.5 seconds with
35 respect to the actual spoken speech. Embedded into a real-time BCI, our approach is capable of providing
36 VAD results with a latency of only 10 ms. *Significance.* To the best of our knowledge, our results show for
37 the first time that speech activity can be predicted purely from unlabeled ECoG signals, a crucial step
38 toward individuals who cannot provide this information anymore due to their neurological condition, such
39 as patients with locked-in syndrome. *Clinical Trial Information.* ClinicalTrials.gov, registration number
40 NCT03567213.

41 [Keywords](#)

42 Brain-Computer Interface, Voice Activity Detection, Electrocorticography

43 [Introduction](#)

44 Several neurological disorders, including amyotrophic lateral sclerosis (ALS), can result in severe paralysis
45 and loss of speech, having devastating effects on the quality of life of affected individuals. Recent
46 advances in implantable Brain-Computer Interfaces (BCIs) have raised hope for the restoration of
47 communication in this clinical population^{1,2} by utilizing neural activity acquired directly from the cerebral
48 cortex to control a neuroprosthetic device that produces text³⁻⁷ or synthesizes speech⁷⁻¹³. Those BCIs are
49 currently trained using supervised learning paradigms where neural activity is mapped onto target
50 representations^{14,15}, such as phonemes or acoustic units, and are therefore dependent on accurate
51 temporal alignments to achieve proper outputs. For this reason, many prior studies in the field have relied
52 on datasets collected from patients who had normal speaking capabilities, such as epilepsy patients^{8,13,16,17}
53 or patients who underwent surgery for glioma removal^{9,11} – datasets where the temporal alignment can
54 be obtained from simultaneous acoustic recordings.

55 In recent years, clinical trials have begun exploring the extent to which approaches previously used in
56 normal speaking subjects can be translated to people in actual need for such a technology^{3,7,18,19}, and while
57 those enrolled clinical-trial participants were speech impaired, their diseases had not yet been progressed

58 into a state of total paralysis that prevented inferring such an alignment. However, in cases where the
59 disease has already progressed to the locked-in syndrome (LIS)^{20,21}, it may not be possible to infer the
60 temporal alignment at all from acoustic data. In pioneering work by Guenther et al.²², a participant living
61 with LIS was able to accurately synthesize vowels continuously using a Kalman filter-based decoding
62 approach with closed-loop neurofeedback. Additionally, more recent work by Chaudhary et al. gave a
63 completely locked-in patient a novel means of communications by spelling sentences using a paradigm
64 that required modulating firing rates with respect to auditory feedback²³.

65 In this study, we make a first step toward acoustic-free model training by assuming that no temporal
66 alignment can be obtained from simultaneous microphone recordings. For this early work, we focus only
67 on localizing and identifying neural activity related to speech processes. Voice Activity Detection (VAD)
68 systems play a crucial role in acoustic speech processing fields, such as automatic speech recognition²⁴ or
69 speaker diarization²⁵, where they may be used in early processing stages to exclude non-speech data when
70 computing acoustic features or embedding vectors. Similarly, many recent BCI studies have also utilized
71 approaches to locate and isolate neural activity related to speech production in their pipelines as an
72 intermediary step to constrain the solution space of speech decoding tasks, both for word recognition^{19,26}
73 and synthesis applications¹⁸. Another application for these neural Voice Activity Detection (nVADs)
74 systems of particular relevance to BCIs is to prevent leakage of speech-related activity into computation
75 of baseline statistics within real-time systems. Decoding performance can degrade over time because the
76 feature space may shift linearly beyond the range expected from the training data. nVAD techniques could
77 help here to determine which parts of the neural data should be considered when updating a running
78 baseline, rather than relying only on a fixed time window containing both speech and non-speech activity.
79 To the best of our knowledge, all previous methods have relied on supervised learning machines trained
80 directly on acoustic ground truth^{18,19,27} or labeled information²⁶ inferred from behavioral cues²⁸ –
81 suggesting that such approaches may not translate to individuals where their disease does not allow
82 vocalization or any observable articulatory movements.

83 Here, we present first results on unsupervised detection of neural voice activity from unlabeled ECoG
84 signals. We did so by setting up an experiment in which a clinical trial participant was instructed to read
85 single words, and where the majority of time for each recording session did not carry speech activity – a
86 design decision we actively exploited to automatically assign identified segments as either speech or non-
87 speech classes. We utilized a graph-based clustering approach²⁹ to find structural patterns with a fixed
88 temporal context in high-gamma activity extracted from ECoG recordings, and used those estimated

89 clustering labels to train a recurrent neural network (RNN). In our evaluation, we first quantified the
90 alignment error between estimated labels from the clustering approach and ground truth acoustic speech
91 information to determine ranges of expected error rates. Next, we compared the performance of our RNN
92 architecture trained on those estimated labels with respect to baseline models previously proposed in the
93 literature trained on VAD labels inferred from acoustic speech. From here, we then inspected how well
94 our model translated to unseen words.

95 Material and Methods

96 Participant and experiment design

97 We conducted an experiment with a clinical trial participant (CC01, male) in his 60s with dysarthria due to
98 ALS, who had been implanted with two ECoG arrays with 64 electrodes each (4-mm center-to-center
99 spacing, 2-mm diameter) covering speech and upper-limb cortical areas (Figure 1a). The participant could
100 speak, but his speaking capabilities were limited, and continuous speech was mostly unintelligible due to
101 his neurological condition^{18,19} (speech was rated with 1 point out of 5 on the ALSFRS-R measure³⁰). In a
102 speech production task, we presented single words on a monitor in front of him and gave instructions to
103 read them out loud. For each trial, the target word was presented for 2 seconds following an inter-trial
104 interval of 3 seconds. Overall, the word pool consisted of 50 words³, and each word was repeated twice
105 in each session. We repeated this experiment across 10 days over a period of 9 weeks. Furthermore, we
106 also collected single word data from a larger word pool of 688 words, which we used to quantify
107 generalization towards unseen words. In this corpus, each word only appeared once, and none appeared
108 in the training data. At the start of each recording day, we conducted a syllable repetition task, which was
109 used for normalizing the neural data. The syllable repetition task was constant across all days to achieve
110 similar statistics for the baseline, in accordance with a prior publication with the same study participant¹⁸.

111 Neural data was digitized using a NeuroPort System (Blackrock Neurotech, Salt Lake City, UT, USA) with a
112 sampling rate of 1 kHz. Audio data was recorded at 48 kHz using an external microphone (BETA[®] 58A,
113 SHURE, Niles, IL). We used BCI2000³¹ for stimulus presentation and for aligning neural and acoustic signals
114 for offline analysis. The clinical trial (ClinicalTrials.gov, NCT03567213) was approved by the Johns Hopkins
115 University Institutional Review Board (IRB) and by the FDA (under an investigational device exemption) to
116 test the safety and preliminary efficacy of a brain-computer interface composed of subdural electrodes
117 and a percutaneous connection to external EEG amplifiers and computers. The participant gave informed

118 consent after being counseled about the nature of the research and implant-related risks, and was
119 implanted with the study device in July 2022.

120 [Cortical mapping](#)

121 The positioning of both subdural ECoG grids was determined via anatomical landmarks from pre-operative
122 structural (MRI) and functional imaging (fMRI). After the surgical implantation of the grids, we conducted
123 a post-operative CT scan, which was co-registered to a pre-operative MRI for verification of the anatomical
124 locations of the two grids. Figure 1a shows a rendering of the participant's brain and the locations of both
125 electrode grids, where the 64 electrodes highlighted in orange were relevant in this study with respect to
126 prior observations¹⁸ about encoded speech activity.

127 [Signal processing and feature extraction](#)

128 We obtained speech-related features from raw ECoG signals by extracting the high-gamma (HG) band,
129 which has shown to track closely the location and timing of speech production neural activation^{32,33} and
130 has been successfully employed in previous studies for speech BCIs^{4,18,19,34–36}.

131 First, we removed all bad channels (19, 38, 48 and 52) based on visual inspection and applied a common
132 average referencing (CAR) filter across each grid independently. Next, we selected the top 64 channels
133 with the strongest activation during overt speech production, identified in a previous study¹⁸ with the
134 same clinical trial participant. We then used a bandpass filter (IIR Butterworth, 4th order) to extract the
135 broadband HG band in the range of 70 - 170 Hz and a notch filter (IIR Butterworth, 4th order) to attenuate
136 the first harmonic of the line noise in the range of 118 - 122 Hz. Finally, for each channel we computed
137 logarithmic power features with respect to a window size of 50 ms and a frame shift of 10 ms. We
138 normalized all features to zero mean unit variance (z-score normalization) with respect to a syllable
139 repetition task conducted at the beginning of each recording day to calibrate the system for day-specific
140 high-gamma changes (see supplementary Figure S1 about the stability of the ECoG signals during the study
141 period). Before using these features for baseline model training, we augmented each frame with a context
142 stacking of 6 consecutive intervals to model temporal dependencies of up to 300 ms in the past. This step
143 was not included in the clustering procedure as the clustering algorithm itself manages a fixed window of
144 past frames to account for the temporal relationships in each cluster.

145 The acoustic data for performance evaluation was collected at 48 kHz, resampled to 16 kHz and
146 segmented into corresponding windows of 50 ms and 10 ms frameshift to match the alignment with the

147 HG features. We verified that no channels had been contaminated with acoustic artifacts by using
148 Roussel's method³⁷. The details of the contamination report are given in supplementary Figure S2.

149 Unsupervised temporal localization of speech production

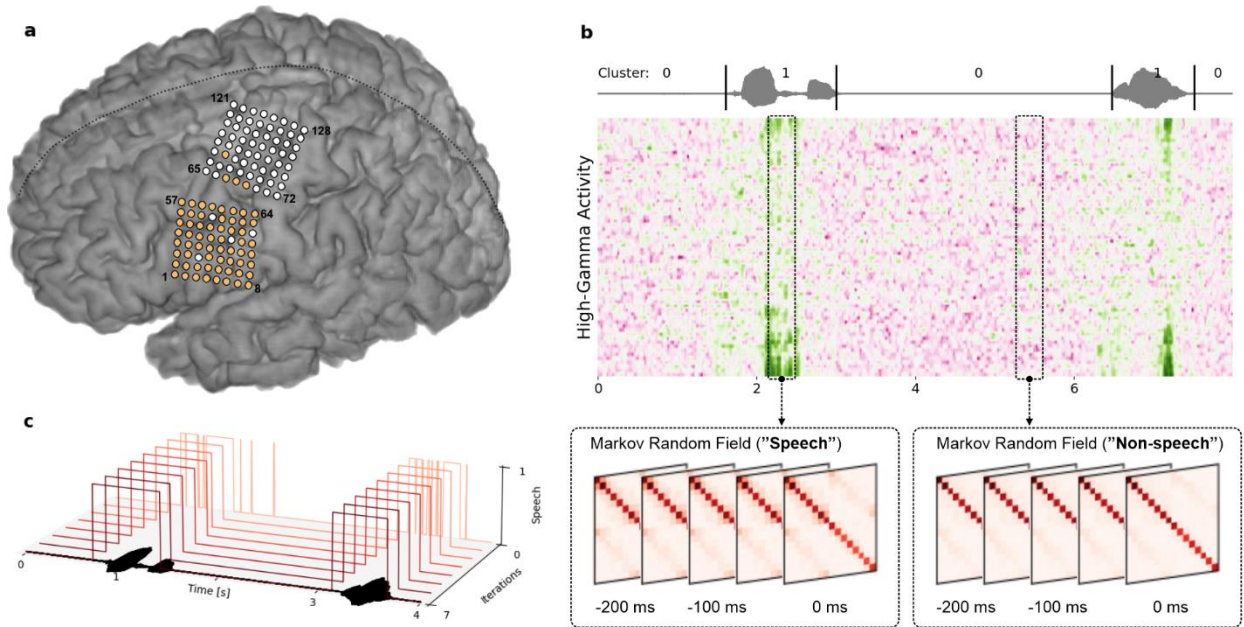
150 To identify speech-associated activity in neural recordings, we adopted a graph-based clustering approach
151 named Toeplitz Inverse Covariance-based Clustering (TICC), specifically designed for discovering common
152 subsequences in multivariate time series data. This unsupervised algorithm defines one Markov Random
153 Field (MRF) per cluster and describes relationships in the form of connections between input features. In
154 our study, these connections would describe dependencies between the neural activity of different
155 electrodes, both with respect to spatial and (potentially) causal temporal patterns.

156 TICC's training procedure is based on an iterative optimization method that employs a variation of the
157 expectation maximization algorithm which first alternates cluster assignments before updating its cluster
158 parameters. Here, the cluster assignment step is based on the path with the minimum cost, obtained
159 using a dynamic programming paradigm. Once this path has been found, the maximization step updates
160 the cluster parameters based on the assigned data points. The training procedure converges when no
161 data points are assigned to a different cluster and are therefore stationary.

162 Besides the number of clusters, the TICC algorithm can be configured with respect to the length of the
163 temporal context and regularization parameters. By specifying multiple layers for the MRFs, data points
164 won't get clustered in isolation but in context to neighboring past observations, allowing it to learn cross-
165 time relationships. Note that temporal layers in the MRFs also obey the Toeplitz constraint to be time-
166 invariant. The regularization parameters β and λ signify the penalty factor for adjacent subsequences
167 being assigned to the same cluster and denote the sparsity level in the MRF's graph structure
168 characterizing each cluster, respectively. A higher β value will result in a greater likelihood of adjacent
169 subsequences being assigned to the same cluster.

170 Figure 1b shows an illustration of the TICC clustering approach. Two MRFs segment the high-gamma
171 activity into speech and non-speech classes. In this example, both MRFs have multiple layers to not only
172 draw insights from spatial characteristics but also capture temporal dynamics of up to 200 ms into the
173 past. The gray waveform at the top has been time-aligned to the neural recordings for visual attribution
174 of the high-gamma activity. Although the clustering assignments do not reveal which clusters belong to
175 speech activity due to their unsupervised nature, we can infer cluster classes based on the length of their
176 subsequences – exploiting the setup of the experiment design. Fig 1c visualizes the clustering process for

177 one recording session. The x-axis represents time and shows a snippet of two trials and the acoustic
178 speech signal as a reference guide. The z-axis shows each iteration from the TICC algorithm until
179 convergence, where found cluster alignments are plotted as curves. The y-axis indicates found speech
180 activity. We based our initial alignments (iteration 0) on clusters found by a Gaussian mixture model, and
181 iteratively optimized those using the TICC algorithm.



182 **Figure 1 | Overview of the experiment setup and clustering approach.** **a** Placement of implanted
183 electrode grids covering speech and upper limb cortical regions. Electrodes highlighted in orange were
184 selected for this study based on previous reported results¹⁸. **b** Illustration of the TICC clustering approach
185 to identify speech and non-speech segments in each trial using one Markov Random Field per cluster. **c**
186 Visualization of the iterative clustering process of the TICC algorithm, starting from an initial alignment
187 derived from Gaussian mixture clustering, until convergence. The acoustic waveform on the x-axis serves
188 as a reference to the found speech clusters.

189 Neural voice activity detection approach

190 We based our nVAD model on the same recurrent neural network architecture from our previous study
191 on synthesizing speech online¹⁸, originally inspired by the work from Zen et al.³⁸ For this binary
192 classification task, all recurrent layers utilize long-short term memory cells³⁹ to learn the temporal
193 dynamics across the individual channels. In total, the network architecture comprises three layers: two
194 LSTM layers with 128 units each and one linear layer with 2 output units, resulting in 231,682 internal
195 weight parameters. We used the cross-entropy loss in conjunction with the softmax activation function

196 to estimate the error between network predictions and target labels during network training, and
197 employed Adam⁴⁰ as our optimizer with a learning rate of 0.0001 and trained the architecture for 20
198 epochs in each fold, while storing the best performing weights in accordance to the minimum validation
199 loss. Network training uses the truncated backpropagation through time (BPTT) algorithm⁴¹ with
200 hyperparameters k_1 and k_2 set to 50 frames of high-gamma activity, respectively, such that the unfolding
201 procedure was limited to 50 frames (0.5 s) and repeated every 50 frames (0.5 ms).

202 Closed-loop system design

203 We built a real-time BCI that communicates directly with BCI2000 about any segments identified as
204 speech. This system was implemented on top of *ezmsg*⁴² – a Python framework that facilitates the
205 development of closed-loop streaming applications by enforcing a software architecture composed of a
206 directed acyclic graph structure. Each node in this graph is responsible for a particular self-contained task,
207 such as computing high-gamma features from raw ECoG signals. We used a network of such nodes to
208 perform tasks that receive ECoG signals, compute features, predict voice activity and communicate results
209 back to BCI2000, including logging functionality between all mentioned nodes for evaluation. In the
210 backend, *ezmsg* utilizes asynchronous coroutines to enable concurrent executions of those tasks. Our
211 closed-loop processing pipeline was capable of producing low-latency feedback as the accumulated
212 computational cost did not exceed the frameshift of 10 ms. Communication with BCI2000 was based on
213 ZeroMQ (ZMQ) as a networking abstraction layer.

214 Results

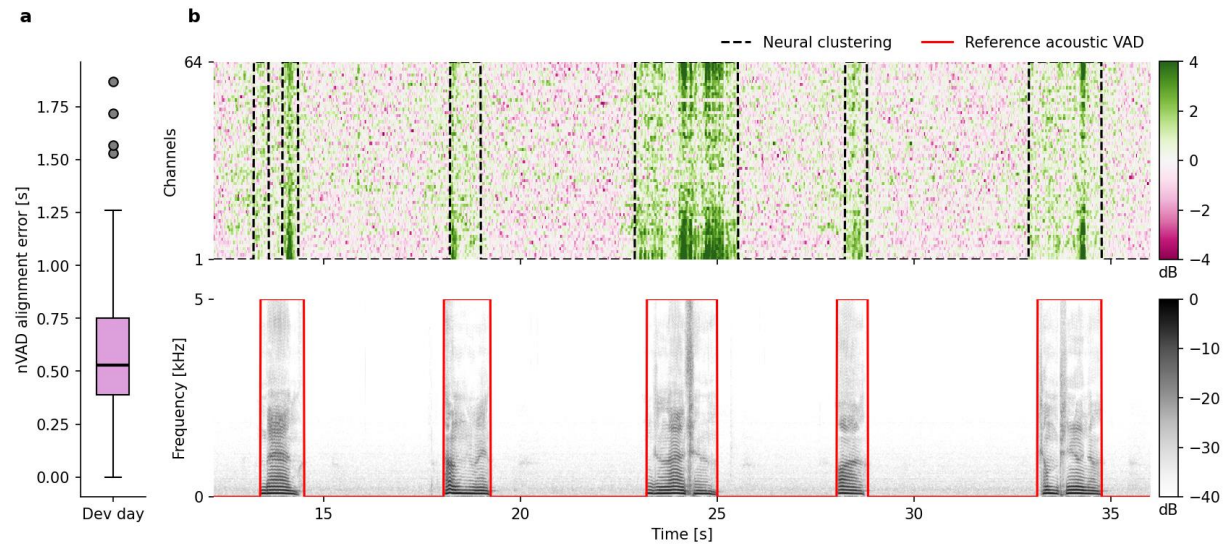
215 Identification of speech segments

216 In this study, we only distinguished between speech and non-speech segments in the neural data, so that
217 all words were summarized in one speech cluster. Another potential approach would be to cluster for
218 each stimulus individually, assuming they were known upfront from the experiment design. However,
219 preliminary analyses suggested that resulting clusters per word find less reliable cluster parameters,
220 potentially converging towards clusters that only identify part of speech segments. We hypothesize that
221 this is related to the inherently smaller amount of data and less variability in the neural activity. When
222 clustering for all words, it is not required to know a specific stimulus or the number of stimuli in advance
223 and is thus also suitable for experiment designs where open questions are asked. We obtained ground
224 truth voice activity information from time-aligned acoustic spectrograms of the microphone recordings
225 which were only used to quantify the accuracy of identified speech segments. We based our evaluation

226 metric on the Levenshtein distance to determine the minimum distance between estimated VAD labels
227 and acoustic VAD ground truth, where all operations for changes were assigned a fixed cost of 10 ms.

228 To infer suitable hyperparameters for the TICC algorithm we utilized ECoG recordings from a single patient
229 with drug-resistant epilepsy (male, between 16-20 years old) who had undergone video-EEG monitoring
230 to localize his seizure onset zone. We particularly chose this data as the implanted ECoG grid covered
231 cortical speech areas similar to our clinical trial participant (see supplementary Figure S3 for details about
232 the grid placement in the epilepsy surgery patient). Note that the electrode grid in the epilepsy surgery
233 patient was implanted in the right hemisphere, yet we were able to measure strong speech-related high
234 gamma activity during speech production. Similar observations have previously been reported in the
235 literature¹³. We ran a grid search across predefined ranges for the β and λ hyperparameters and selected
236 those which achieved lowest alignment errors with respect to ground truth voice activity of the epilepsy
237 patient, leading to a hyperparameter configuration of $\beta = 50$ and $\lambda = 11e^{-4}$.

238 Our results are summarized in Figure 2a. On a held-out day used to report intermediate results from the
239 clustering algorithm (from now on referred to as development set), we achieved a median alignment error
240 of 530 ms per trial, while 75% of the trials were below 752 ms (average speech duration: 1.2 s). In 8 out
241 of 400 trials, speech could not be detected through the clustering approach and, additionally, 10 trials
242 resulted in alignment errors above the average speech duration of 1.2 s. Figure 2b shows an excerpt of
243 the first 5 trials of the first day in the training set for visual inspection. The top panel visualizes high-gamma
244 activity and how frames have been clustered after applying the TICC algorithm with the same
245 configuration of hyperparameters obtained from the epilepsy patients data. The bottom panel shows the
246 time-aligned speech spectrogram and ground truth VAD information based on the acoustic signal. Overall,
247 the clustering approach can identify consecutive segments of spoken speech reliably in the majority of
248 the cases, leading to labels that can be utilized to train a supervised model that predicts speech activity
249 for an incoming stream of high-gamma frames without calculating the minimum alignment path using
250 dynamic programming strategies.



251

252 **Figure 2 | Comparison between VAD labels estimated from acoustic and high-gamma representations.**

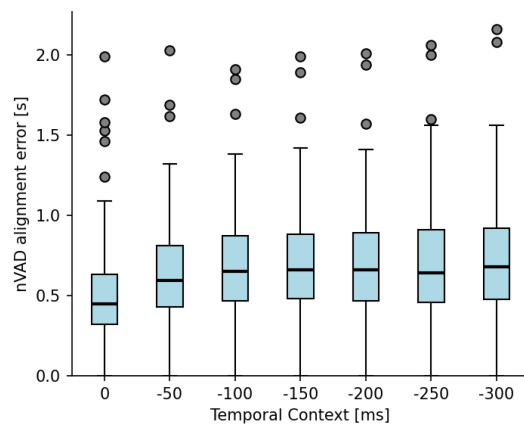
253 **a** Minimum alignment error computed via Levenshtein distance between neural speech clusters and
254 acoustic reference VAD using the development set ($n = 400$ trials). Box indicates boundaries between
255 quartiles Q1 and Q3, and whiskers represent range of data within 1.5 times the interquartile range.
256 Outliers correspond to trials for which no speech clusters could be found from the neural activity. **b** Visual
257 example of the first 5 trials from the first day in the training set. Top panel shows estimated speech
258 clusters using the TICC algorithm (dashed black line) on high-gamma features and bottom panel the
259 corresponding time-aligned speech spectrogram from the acoustics with reference VAD (solid red line).

260 [Temporal context provides less accurate speech clusters](#)

261 Next, we analyzed if nVAD labels can be more accurately determined by including causal temporal
262 contextual information. Here, we adapted the TICC algorithm to avoid repetitive information from the 40
263 ms overlaps in the feature extraction pipeline by adding a dilation hyperparameter indicating the spacing
264 between consecutive high-gamma frames. In accordance with Soroush et al.²⁷, we investigated temporal
265 dynamics up to 300 ms into the past. MRFs with only one layer correspond to no context information,
266 with five layers up to 200 ms into the past (as represented in Figure 1b), and with 7 layers of up to 300
267 ms, where each additional layer introduces a dilation of 5 frames to avoid repetitive information from the
268 40 ms overlap in the feature extraction pipeline.

269 Similar to Figure 2a, we report our observations on the development set and used the minimum distance
270 between estimated VAD labels and ground truth labels calculated on the speech spectrogram as the error
271 metric, again with a cost of 10 ms per off-diagonal step in the alignment matrix. Figure 3 visualizes our

272 results in the form of boxplots. We found that the median alignment error increased as more temporal
273 context information was captured in each feature vector. We hypothesize that this trend stemmed from
274 the growing number of features enabling more complex relationships in the spatio-temporal connections,
275 which were inadequately supported by the limited amount of data, leading to increasingly inaccurate
276 cluster parameters. We observed similar results with respect to the data used to determine appropriate
277 hyperparameter choices; therefore, all further analyses were conducted with only one-layer MRFs.

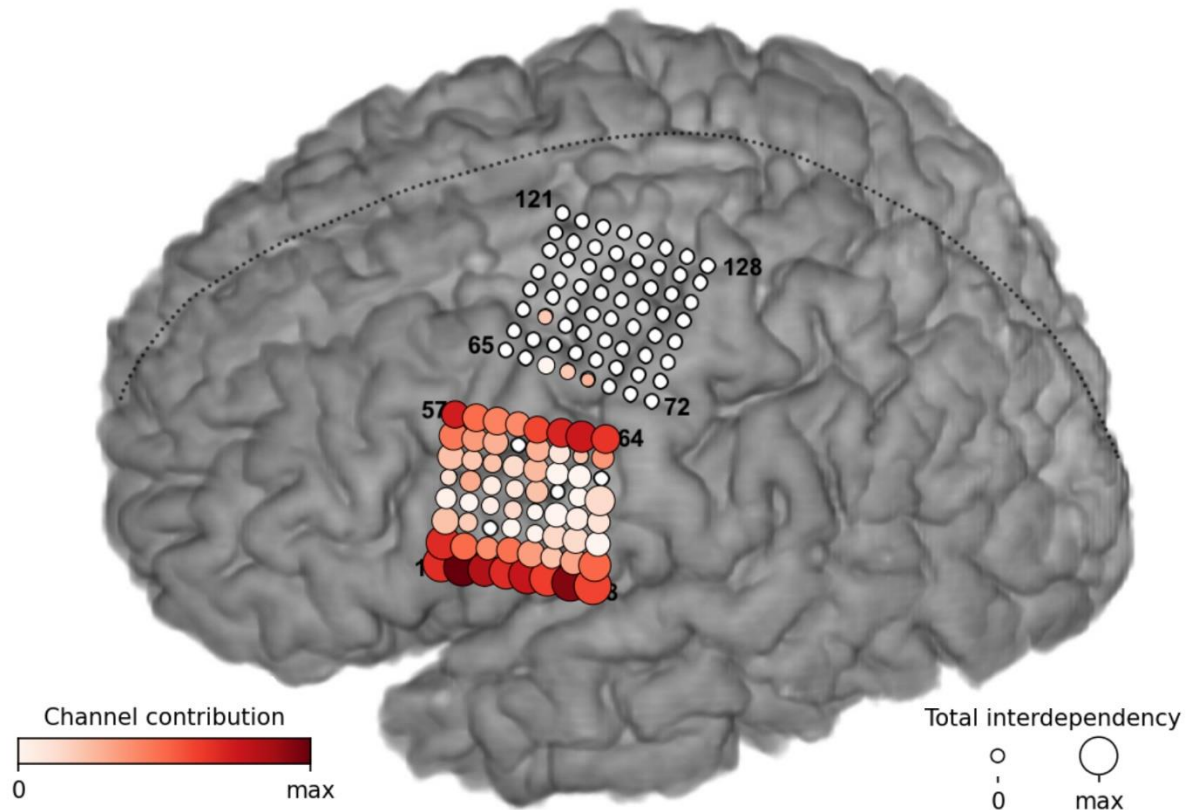


278 **Figure 3 | Adding temporal context leads to less accurate nVAD labels.** Trend of inaccuracies for
279 estimated nVAD labels increases with including context information. No context information and 300 ms
280 into the past correspond to MRF's with number of layers of 1 and 7, respectively. Evaluation was
281 conducted on the development set to report the impact of including temporal context, however, we based
282 our decision on using only one layer in the MRFs on the lowest error score obtained by the TICC algorithm
283 with respect to the data from the epilepsy surgery patient.

284 [Cluster parameters suggest consistent task-specific activity in motor cortices](#)

285 The graphical dependency structures underlying cluster representations allow learned relationships to be
286 interpreted and pinpointed to cortical areas known to elicit activity during speech – enabling us to verify
287 that proper representations have been learned. We analyzed the differences between both speech and
288 non-speech MRFs to reveal which connections between electrodes contribute to what extent to the
289 decision-making process. Our findings are visualized in Figure 4. Each circle on the brain plot belongs to
290 one channel. The color of the circle represents how much a particular channel contributed in the decision-
291 making process of the clustering assignment and the size indicates the total sum of the interdependencies
292 between channels. The plot reveals that the differences in high-gamma activity features from electrode
293 channels located in vM1 and dM1 were predominantly used to discriminate between speech and non-

294 speech clusters in the TICC algorithm. Both of these cortical areas have already shown speech activity to
295 various degrees in our prior publication¹⁸. Moreover, the plot suggest that the algorithm focused on a
296 rather smaller subset of electrodes compared to our prior publication on synthesizing keywords where
297 the supervised nVAD model based its decision on a much broader network of electrodes across motor,
298 premotor and somatosensory cortices. We hypothesize that this is related to the different machine
299 learning approaches (a recurrent neural network compared to the TICC algorithm), the increased number
300 of word stimuli (50 stimuli instead of 6) and the variability in the data as some words in the 50-word
301 corpus are longer and more effortful to articulate.



302 **Figure 4 | Cluster assignments mainly driven by differences in inter-electrode connections in vM1 and**
303 **dM1.** Visualization of the differences in the found MRF structures between both speech and non-speech
304 clusters. The color coding of the circles represents electrode contributions, while the size indicates the
305 strength of inter-channel dependencies. These relationships show that the TICC algorithm focused
306 primarily on spatial high-gamma activity patterns between electrodes in vM1 and dM1 when deciding
307 which cluster to assign.

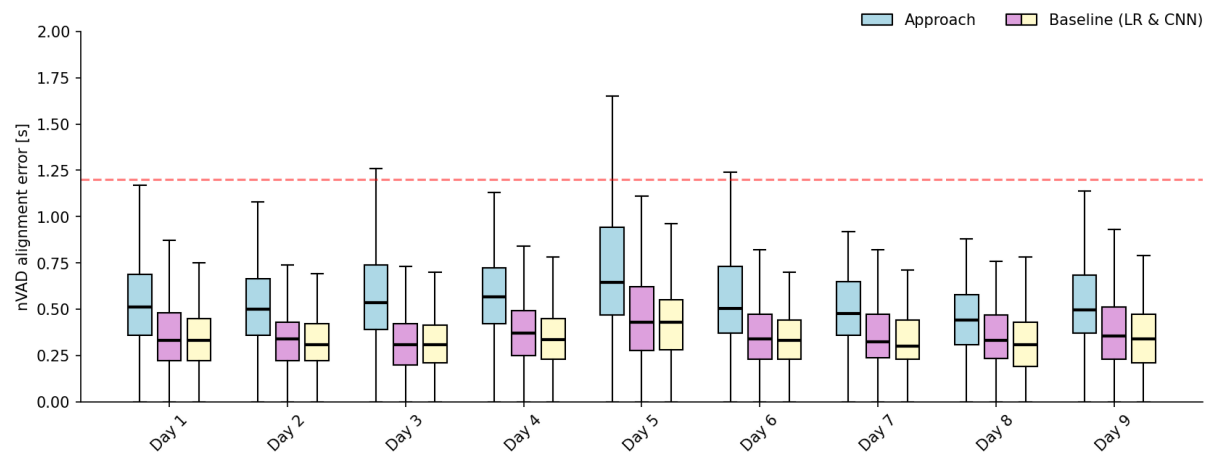
308 Predicting speech from neural activity

309 We evaluated our proposed approach using a leave-one-day-out cross-validation method to quantify
310 model performance across multiple days. Moreover, this prevented day-specific information from the
311 testing days leaking into the training set which may wrongfully bias generalization. We compared our
312 approach trained on the estimated labels from the TICC algorithm against two other methods previously
313 reported in the literature²⁷, namely logistic regression (LR) and a LeNet-style convolutional neural network
314 (CNN)⁴³, both trained on ground truth information acquired from the acoustic speech spectrogram. For
315 these baseline models, we followed the corresponding study by Soroush et. al.²⁷ and used context stacking
316 up to 300 ms into the past, with no additional sequence modeling techniques that would consider outputs
317 from previous time steps.

318 Our results from the cross-validation are summarized in Figure 5. For each day, we report the alignment
319 errors for our approach (light blue) and the two baseline models (pink and yellow) as boxplots (samples
320 per day: $n_1 = 306$, $n_2 = 204$, $n_3 = 204$, $n_4 = 204$, $n_5 = 204$, $n_6 = 204$, $n_7 = 204$, $n_8 = 102$, and $n_9 = 204$). The
321 dashed red line indicates the average speaking duration of 1.2 s per prompted word from our participant.
322 Across all days from the study period, we observed median error scores between 440 and 645 ms, where
323 50% of the trial-based errors were in the range of 380 to 710 ms. Furthermore, we also observed that in
324 5.2% of the trials (96 out of 1836 trials) our model was not capable of detecting speech instances at all or
325 made prediction errors that exceeded the average speaking duration. We excluded those outliers in Figure
326 5.

327 Regarding the baseline methods, the CNN model achieved overall the lowest alignment errors with
328 median scores between 300 and 430 ms across days, where 50% of the trials deviated between 220 and
329 450 ms from the ground truth acoustics. For the logistic regression, the alignment errors were slightly
330 higher between 310 and 430 ms in median, which was in line with previous findings on sEEG data²⁷. Here,
331 50% of the trials had alignment errors between 230 and 490 ms.

332 Although the results from our approach were not on par with baseline models trained on ground truth
333 VAD information, we observed that our approach was still capable of detecting the majority of spoken
334 speech, up to 77% per day, and on average 70% across days. This would be particularly useful for filtering
335 out speech frames during online computations to obtain normalization statistics based on streaming
336 neural activity.



337

338 **Figure 5 | Cross validation results regarding the proposed approach and baseline models.** Alignment
339 errors are reported with respect to the specific held-out day in each fold. Box plots indicate that our
340 approach achieves consistently higher error rates in the range of 140 and 215 ms than models trained on
341 ground truth VAD information.

342 [Generalizability towards unseen words](#)

343 Next, we analyzed the applicability to spoken words beyond our training corpus of 50-word stimuli to
344 quantify generalization. We recorded an additional corpus of 688 words (each word was only repeated
345 once) across 7 sessions on one particular day (outside of the training days) and computed the mean
346 alignment errors for all trials. The average speaking duration regarding of unseen words was 1.3 s per
347 word. Our results do not show any substantial deviations from those word stimuli that were present in
348 the training corpus. The median alignment errors were between 446 and 490 ms, with 50% of the trials
349 occurring in the 340 and 650 ms range, suggesting that this approach is also applicable to unseen word
350 stimuli.

351 [Discussion](#)

352 Here, we demonstrate a BCI that is capable of identifying speech activity in real-time from ECoG signals
353 recorded from speech-related cortical areas in a clinical trial participant living with ALS. Prior studies
354 reporting on voice activity detection from neural activity have relied on ground truth acoustic speech
355 information to train predictive models – a major challenge when translating such findings to paralyzed
356 individuals who have lost their ability to speak. Our approach utilizes a graph-based clustering technique
357 to localize consecutive segments in the neural data related to speech production. We designed an

358 experiment paradigm that can infer which clusters most likely belong to speech activity based on their
359 clustering lengths. By training a recurrent neural network on these estimated alignments, we were able
360 to identify the majority of speech activity in more than 92% of the trials.

361 While the performance of our approach was not on par with baseline models trained on ground truth
362 acoustic speech information, it would not be reasonable to expect equivalent or better performance in
363 the absence of ground truth. The timing and magnitude of muscular contractions preparing for and
364 executing phonation and articulation do not have a one-to-one correspondence with the timing and
365 magnitude of the acoustic waveform produced by speech, which serves as the ground truth for VAD.
366 Consequently, the timing and magnitude of neural activity in sensorimotor cortex, which form the basis
367 for nVAD, are not expected to be perfectly aligned with spoken acoustics. Moreover, while the signal-to-
368 noise ratio of ECoG high-gamma power modulation has proven sufficient for decoding speech, it is
369 nevertheless non-stationary and dependent on imperfect estimates of its noise floor during non-speech
370 segments, derived here from a separate session with cued speech segments. In spite of these challenges
371 for nVAD, we found that our approach could detect the majority of speech. Analyses on seen and unseen
372 word stimuli revealed that recall scores of approximately 78% could be achieved, compared to 89% from
373 the CNN baseline models. While our current approach was not capable of always isolating each spoken
374 word in its own unique segment, additional postprocessing strategies may help prevent such behavior.
375 Such strategies have been used in the past to correct misclassified frames based on a fixed window of
376 predictions⁴⁴.

377 By interpreting and comparing cluster parameters, we found that assignments were mainly driven by
378 differences of neural activity in a subset of the electrodes in the vM1 and dM1 cortical regions and their
379 interconnections. Even though many more electrodes show high-gamma activations during overt speech
380 production, the clustering approach converged to similar weights and interconnection weights for both
381 speech and non-speech MRFs on those electrodes. One explanation of this behavior might lie in the high-
382 gamma activity variability across word stimuli, and that the TICC algorithm identified those less reliably
383 when making the binary assignment.

384 A limitation of our study is that our participant was still able to speak, albeit with significant dysarthria
385 and poor intelligibility. Thus, it remains to be seen if our approach translates to patients who are incapable
386 of producing audible speech. In this study, we focused intentionally on a patient who could still speak so
387 that we could compare the performance of our approach with ground truth speech acoustics and to
388 estimate the extent of alignment errors – which would not have been possible if speech had been absent.

389 In this pilot study, we addressed the open challenge of training a BCI that identifies speech without having
390 time-aligned neural and acoustic data. Our results show that a graph-based clustering approach can
391 identify segments of spoken speech in neural recordings with median alignment mismatches below 500
392 ms. Despite this inaccuracy, we were able to train VAD models and deploy them in a real-time streaming
393 scenario to predict speech activity online. The error rate may be small enough for practical application.
394 We believe this would be particularly useful for avoiding the inclusion of speech frames when calculating
395 baseline neural activity during non-speech segments and for real-time gating of speech decoders in
396 speech BCIs, including brain-to-text and brain-to-speech applications. Moreover, our approach could also
397 benefit BCI systems by acting as a switch to toggle on the decoder when the user generates silent speech,
398 and toggle off after some time of silence. This would prevent undesired random speech decoding when
399 the user is doing other tasks that somehow affect motor activity. Future work is necessary to determine
400 whether our approach is equally effective for individuals who can no longer produce audible speech.

401 Code availability

402 All source code supporting this study will be made publicly available on
403 <https://github.com/cronelab/corticom-neural-vad> upon acceptance of the manuscript. Moreover, the
404 repository also comes with a bash script which can be used to replicate all steps done in this study,
405 including rendering the figures and running the real-time BCI on streamed signals.

406 Data availability

407 All data supporting this study will be made publicly available on www.osf.io upon acceptance of the
408 manuscript. Neural recordings are prepared in the MATLAB file format version 5, where time-aligned
409 anonymized acoustic speech is stored in the wav file format.

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503 Author Contributions

504 M.A. and N.C. wrote the manuscript. M.A., S.J., S.L., Q.R. and D.C. analyzed the data. M.A. S.L. and Q.R.
505 collected the data. M.A., S.J. and G.M. implemented the code for model training and system design. M.A.
506 and S.J. made the visualizations. C.G., K.R., L.C. and N.M. conducted the medical procedure. F.T. handled
507 the regulatory aspects. N.C., N.R. and M.F. supervised the study and the conceptualization. All authors
508 reviewed and revised the manuscript.

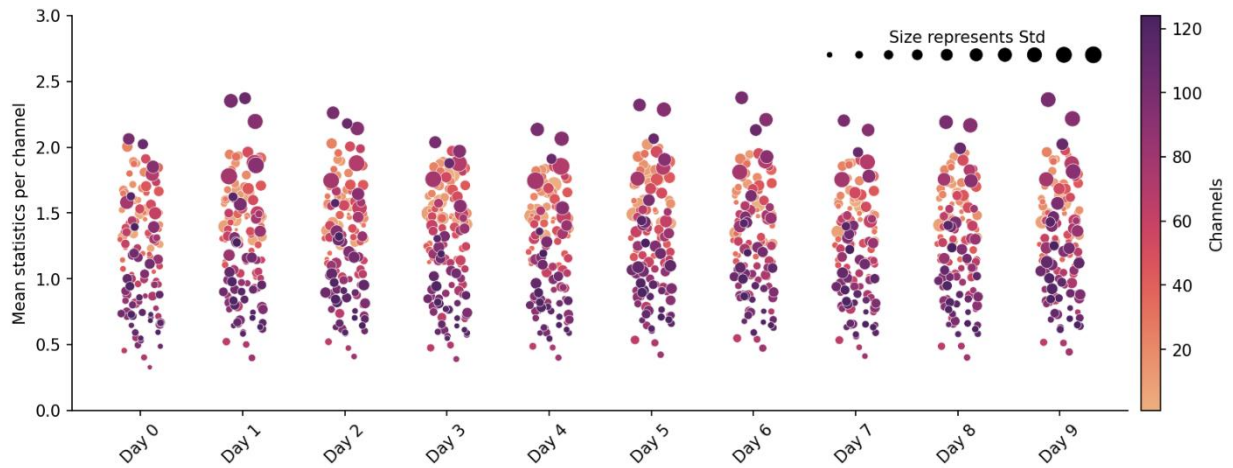
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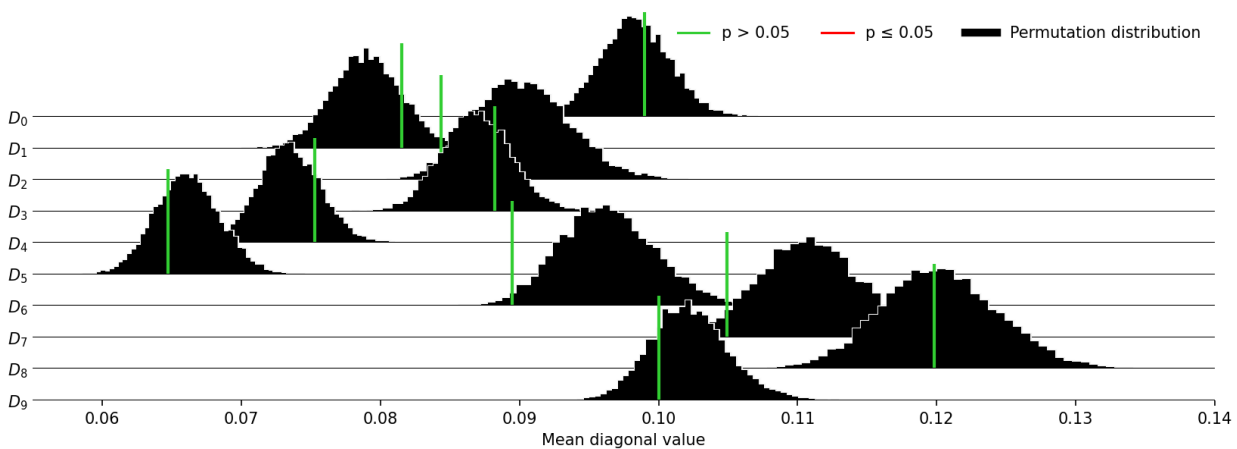
514 Competing Interests

515 The authors declare that they have no competing interests.

516 Supplementary Material

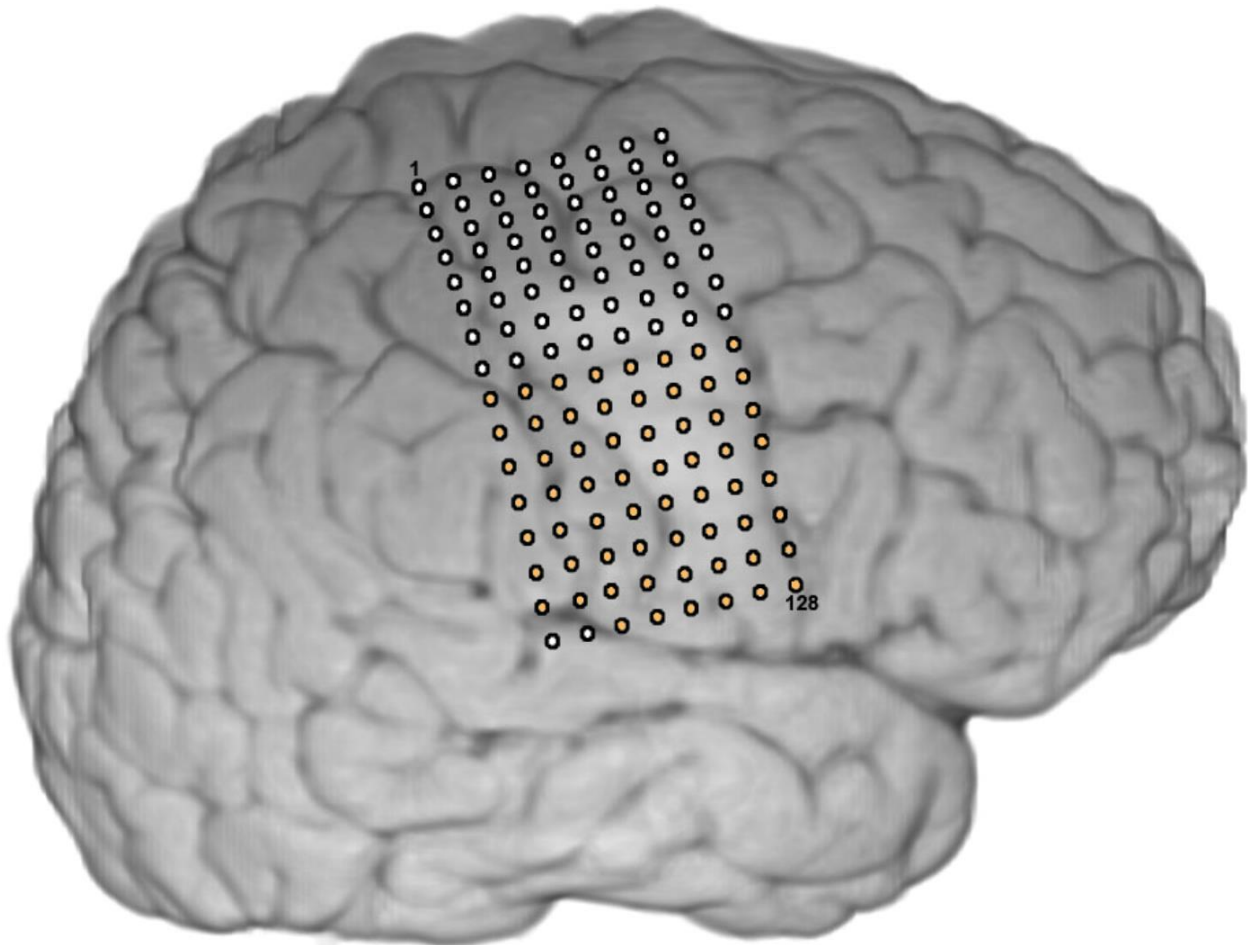


517 **Supplementary Figure S1 | High-gamma ECoG activity remained stable across all study days.** Similar
518 structures can be observed for the majority of the channels, indicating that those channels had
519 comparable activity values and were stable during the study period. Randomized channel shifts on the x-
520 axis were conducted using the same noise profile for all days to only encode relevant information with
521 respect to the y-axis, size and color. Bad channels 19, 38, 48 and 52 were omitted in this plot.



522
523 **Supplementary Figure S2 | Summary of the acoustic contamination report.** All recordings within our
524 development set (D₀) and the days for the open-loop recordings (D₁ - D₉) were checked for acoustic
525 contamination by using Roussel's method³⁷. Each histogram visualizes for one day the distribution of mean
526 diagonal values from permutated contamination matrices (N=10,000 permutations). The vertical-colored
527 bars represent the actual mean diagonal value of the contamination index. The statistical criterion for
528 rejecting the null hypothesis is either displayed in green (p > 0.05) or red (p ≤ 0.05) indicating that the

529 neural signals have been acoustically contaminated. We observed in one channel contamination artefacts
530 for one day (day 7) and replaced those high-gamma values with mean activity from neighboring channels.
531 After this step, no acoustic contamination was present anymore. All recordings from the closed-loop block
532 were omitted here as they have not been used for model training.



533
534 **Supplementary Figure S3 | ECoG array placement in an epilepsy patient to infer a suitable**
535 **hyperparameter configuration.** We determined appropriate hyperparameters for the TICC algorithm by
536 using data recorded from an epilepsy patient implanted with a 128 channel ECoG grid covering similar
537 speech areas than our clinical trial participant. We selected the bottom 62 channels (2 electrodes covering
538 superior temporal gyrus were excluded) to roughly match similar areas than our clinical trial participant.
539 Although the ECoG grid in this patient was implanted on the right hemisphere we observed strong high-
540 gamma activity during speech production, supporting similar observations previously reported in the
541 literature¹³.