



BACKGROUND

- Individuals with Autism Spectrum Disorder (ASD) often have difficulties with various aspects of language processing, including *semantic processing*: the ability to receive a stimulus and apply meaning based on previously stored knowledge [1]
- The *N400 Effect*, an established ERP component elicited upon recognition of a disruption in semantics, occurs from approximately 300-500 ms [2] and is generally reduced or absent in response to linguistic stimuli in ASD [3], suggesting impairments in semantic processing
- This difficulty with semantic processing could arise from an impairment in the language system (**Fig. 1a**). However, findings of intact word decoding and hyperlexia [4] in ASD rule out this scenario.
- Alternatively, the semantic processing system could be impaired in ASD (**Fig. 1b**). However, findings of intact semantic processing of non-linguistic stimuli like pictures and environmental sounds [3,5] rule out this scenario.
- A final possibility is that the connections between the language and semantic systems are impaired (**Fig. 1c**). This has been proposed by previous neuroimaging work finding atypical structural connectivity in ASD brains, notably in left fronto-parietal areas (linked to semantic/language processing) and their connections [6]
- However, the temporal aspects of this underconnectivity remain unclear.
 - If impaired communication occurs in pre-semantic time windows, underconnectivity should be apparent before N400 onset.
 - If impaired communication occurs at post-lexical integration steps, underconnectivity should be apparent after N400 offset.
- EEG coherence* time-locks changes in coherence (the synchronicity of oscillations) to specific events or changes in stimuli, recording changes in the timecourse of neural activation and connectivity [7]
- Identifying the temporal focus of underconnectivity during lexico-semantic processing could provide important insights into which particular aspect of neural communication is going awry.

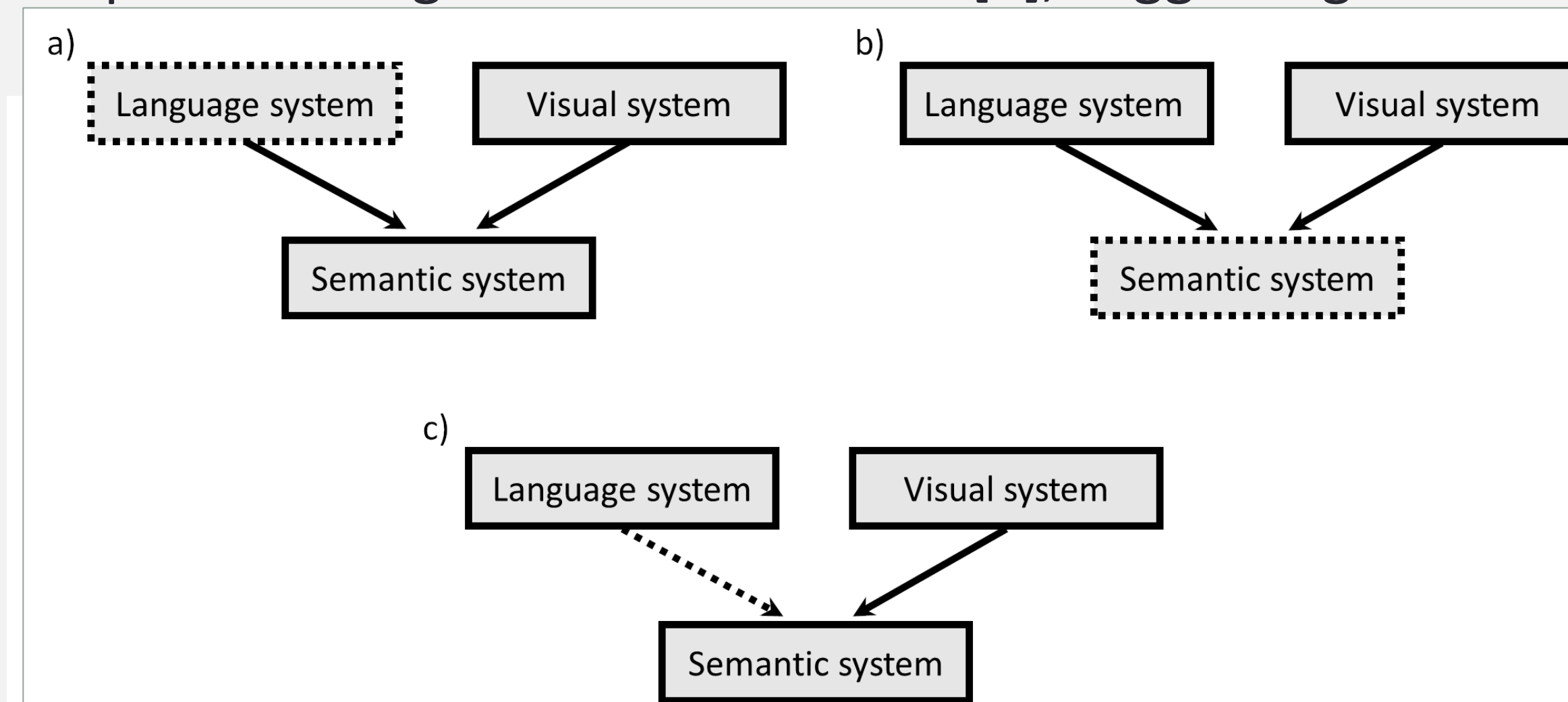


Fig. 1: Theoretical model of lexico-semantic impairments in ASD

RESULTS

- Early after the first and second stimuli (100-300 ms), the TD group showed large peaks in theta coherence at fronto-parietal connections (F3-P3 and F4-P4).
- After both word stimuli, the ASD group showed trends of reduced theta coherence at the left fronto-parietal pair (F3-P3; all p 's<0.07) but not the right fronto-parietal pair (F4-P4; all p 's>0.86); no group differences for pictures.

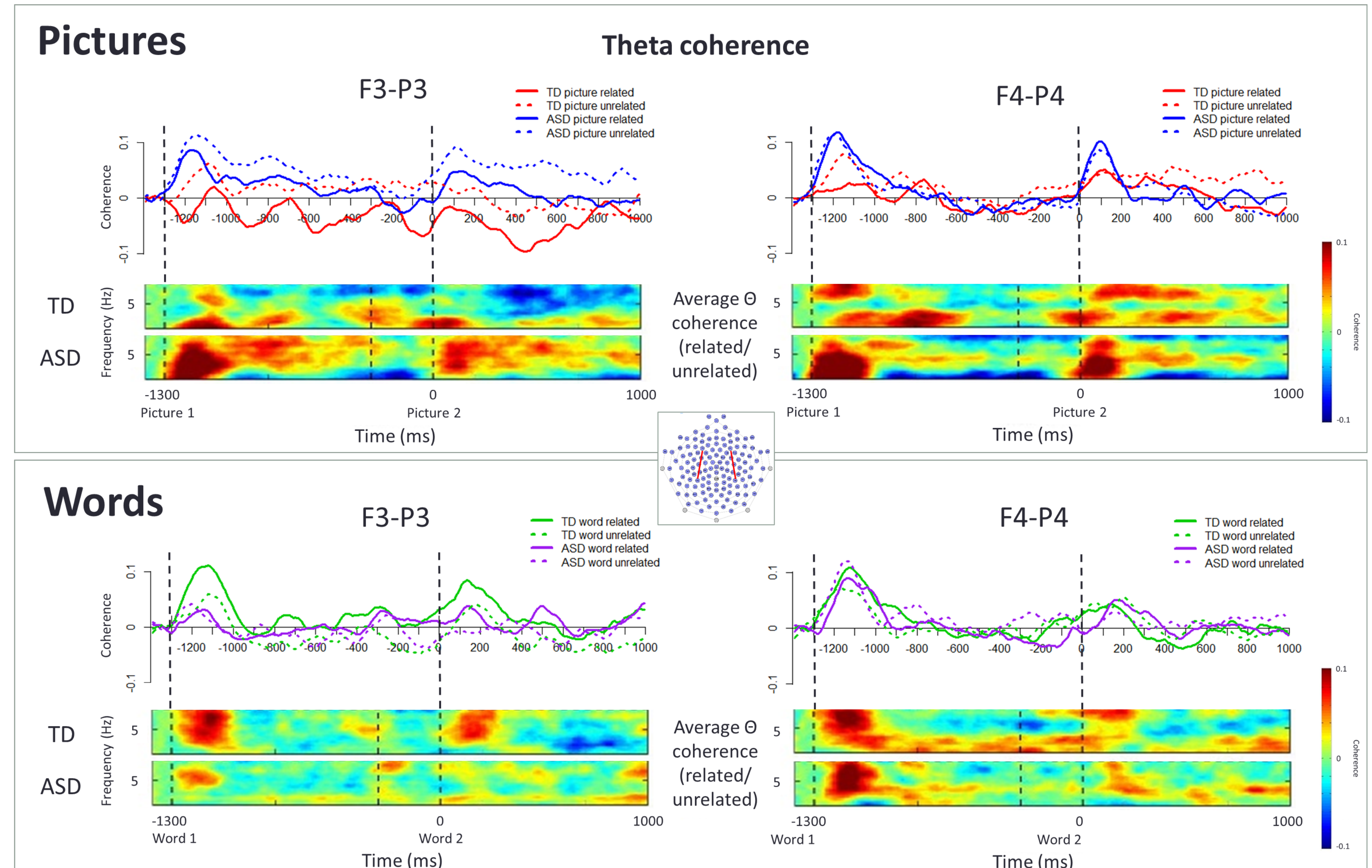


Fig. 3: Theta coherence at left fronto-parietal (F3-P3) and right fronto-parietal (F4-P4) electrode pairs. Top: line graphs of averaged theta coherence over the entire frequency band in each group and condition. Bottom: full frequency distribution in the theta band.

CONCLUSIONS

- The observation of group differences in early (100-300ms) peaks in theta coherence at left fronto-parietal connections, in the language domain only, supports our prediction that neural connectivity during lexico-semantic processing is selectively impaired.
 - This underconnectivity was absent in the analogous right-hemisphere pair, replicating prior findings that underconnectivity in ASD is more prominent in left fronto-parietal areas [6].
- Our data identify a clear temporal locus of underconnectivity: 100-300 ms after stimulus presentation, clearly before the onset of the N400 ERP component.
- In accordance with a recent model [9], early increases in theta coherence could represent semantic information flowing from temporal to frontal areas (rising phase of the N400).
- Future directions include an extension to fMRI, where we plan to replicate the paradigm to confirm that underconnectivity is centered around the left-hemisphere connections.

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METHODS

The data used here is the same as that reported in our prior study examining the N400 ERP component in this paradigm [5]. The participants and experimental methods described here can also be found in that publication.

Participants

- 20 adults with ASD (M = 28 years); 20 typically-developing (TD) participants (M = 25 years).
- Groups matched on age ($p=0.34$). Groups differed on receptive vocabulary ($p=0.07$) and verbal/non-verbal IQ ($p<0.05$); these were included as covariates in analyses.

Stimuli and Procedure

- Participants viewed pairs of pictures or pairs of words that were semantically related or unrelated (100 pairs per trial type).
- Participants monitored for “catch trials” (16% of stimuli) in which a smiley face (picture blocks) or consonant string (word blocks) were presented, and were instructed to hit a button when they saw the catch stimulus. Catch stimuli were not included in analyses.

EEG Data Acquisition, Preprocessing, and Analysis

- EEG data recorded at 500 Hz using a 128-channel Geodesics Sensor net and NetStation 5.3.
- Data bandpass filtered from 0.1-50 Hz and segmented into epochs time-locked to the onset of the first stimulus.
- Time-frequency analysis of theta band (3.5-7 Hz); Morlet wavelet of 2 cycles with expanding factor of 0.5 and Hanning taper.
- Coherence calculated for 12 intrahemispheric electrode pairs based on 9 electrodes taken from the 10-20 distribution system [8]; at 393 frequencies from 2-50 Hz (approximately every 0.1 Hz) and at 300 time points from -242 ms to 2440 ms around first stimulus
- Theta frequency band defined as 3.5-7.5 Hz

	Pre-trial fixation 200 ms	Stimulus 1 1000 ms	Inter-stimulus fixation 200 ms	Stimulus 2 1000 ms	Inter-trial interval 1000-1400 ms
Picture, related	+		+		+
Picture, unrelated	+		+		+
Picture, catch trial	+		+		+
Word, related	+	cat	+	dog	+
Word, unrelated	+	frog	+	clock	+
Word, catch trial	+	XPLKW	+	bread	+

Fig. 2: Examples of stimuli from the experimental paradigm