1 Title page

- 2 **Title:** Automated classification of cognitive decline and probable Alzheimer's dementia across
- 3 multiple speech and language domains
- 4 **Running title:** Detect pAD across subsections & language domains
- 5 Names of the authors: Rui He^a, Kayla Chapin^a, Jalal Al-Tamimi PhD^b, Núria Bel PhD^a, Marta
- 6 Marquié MD, PhD^{c,d}, Maitee Rosende-Roca MD^c, Vanesa Pytel MD, PhD^c, Juan Pablo Tartari
- 7 MD^c, Montse Alegret PhD^{c,d}, Angela Sanabria PhD^{c,d}, Agustín Ruiz MD, PhD^{c,d}, Mercè Boada
- 8 MD, PhD^{c,d}, Sergi Valero PhD^{c,d}, Wolfram Hinzen PhD^{a,e}

9 Affiliations and addresses of the authors:

- 10 ^a Department of Translation & Language Sciences, Universitat Pompeu Fabra, Carrer Roc
- 11 Boronat, 138, 08018 Barcelona, Spain.
- ^bLaboratoire de Linguistique Formelle (LLF), CNRS, Université Paris Cité, Bâtiment Olympe
- 13 de Gouges, 5ème étage. 8, Rue Albert Einstein 75013 Paris.
- 14 °Ace Alzheimer Center Barcelona, Universitat Internacional de Catalunya, C/Gran Via de
- 15 Carles III, 85 bis, 08028 Barcelona, Spain.
- 16 ^dNetworking Research Center on Neurodegenerative Diseases (CIBERNED), Instituto de
- 17 Salud Carlos III, Madrid, Spain.
- 18 ^eIntitut Català de Recerca i Estudis Avançats (ICREA), Barcelona, Spain, Passeig de Lluís
- 19 Companys, 23, 08010 Barcelona, Spain.
- 20 Correspondence: Rui He, Department of Translation & Language Sciences, Universitat
- 21 Pompeu Fabra, Carrer Roc Boronat, 138, Barcelona 08018, Spain, rui.he@upf.edu,
- 22 @RuiHe76864182

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39 Abstract

40 *Background:* Decline in language has emerged as a new potential biomarker for the early 41 detection of Alzheimer's disease (AD). It remains unclear how sensitive language measures 42 are across different tasks, language domains, and languages, and to what extent changes can be 43 reliably detected in early stages such as Subjective Cognitive Decline (SCD) and Mild 44 Cognitive Impairment (MCI).

45 Methods: Using a scene construction task for speech elicitation in a new Spanish/Catalan speaking cohort (n = 119), we automatically extracted features across seven domains, three 46 47 acoustic (spectral, cepstral, voice quality), one prosodic, and three from text (morpho-lexical, 48 semantic syntactic). They were forwarded to a random forest classifier to evaluate the discriminability of participants with probable AD dementia (pAD), amnestic and non-amnestic 49 50 MCI, SCD, and cognitively healthy controls. Repeated measure ANOVAs and paired-sample Wilcoxon sign-ranked test were used to assess whether and how performance differs 51 52 significantly across groups and linguistic domains.

Results: The performance scores of the machine learning classifier were generally satisfactorily high, with the highest scores over .9. Model performance was significantly different for linguistic domains (p < .001), and speech vs. text (p = .043), with speech features outperforming textual features, and voice quality performing best. High diagnostic classification accuracies were seen even within both cognitively healthy (controls vs. SCD) and MCI (amnestic and non-amnestic) groups.

59 *Conclusions:* Speech-based machine learning is powerful in detecting cognitive decline and
 60 pAD across a range of different feature domains, though important differences exist between

- 61 these domains as well.
- 62 Keywords: Machine Learning; probable Alzheimer's Disease; Connected Speech

64 Introduction

Although Alzheimer's disease (AD) is one of the leading causes of death in older adults, there 65 66 are no drugs in clinical practice that can cure or prevent the disease (Mossello & Ballini, 2012; World Health Organization, 2020). In this regard and the context of global aging, efficient and 67 accessible approaches to predicting AD risk at earlier stages have been widely sought, 68 69 including in Mild Cognitive Impairment (MCI) and even preclinically, in individuals with 70 Subjective Cognitive Decline (SCD) (Rabin et al., 2017). Traditional methods for AD detection 71 suffer from a number of limitations, including invasiveness and high cost (e.g., lumbar puncture, 72 neuroimaging markers), or low specificity (e.g., Mini-Mental State Examination (MMSE) and 73 Clinical Dementia Rating (CDR)). In this context, language has emerged as a new and potentially promising biomarker for detecting AD at very early stages, and developing with 74 75 disease progression (Ahmed et al., 2013; Uretsky et al., 2021). It is noteworthy that people with 76 SCD, by definition, know that they are complaining, and that such awareness may impact on 77 speech parameters, beyond potential organic factors relating to their elevated risk of AD. 78 Automation of speech and language analysis is rapidly advancing, and theoretical models 79 support the integration of language and memory networks in the brain, pointing to shared underlying neural mechanisms (Hagoort, 2019; Roger et al., 2022). The integration of these 80 81 methods with other signals from behavior and/or with biological markers such as blood may 82 prove essential for advancing on inexpensive, widely available, and robust markers of early 83 disease progression in AD.

Automated approaches from natural language processing (NLP) and machine learning have already shown strong capability in predicting AD, and even MCI (de la Fuente Garcia et

86 al., 2020). Paralinguistic measures extracted from speech directly, such as mathematical properties of sound waves, have also shown impressive power in identifying AD (Chen et al., 87 88 2021; Haider et al., 2020; Sarawgi et al., 2020). Some studies transcribed audio into texts, either 89 by hand or machine, and extracted features from texts, including lexical, syntactic, and N-gram features (Orimaye et al., 2017; Qiao et al., 2021; C. Thomas et al., 2005). Unlike these 90 91 traditional feature engineering approaches, transfer learning based on pretrained language 92 models, such as bidirectional encoder representations from transformers (BERT), encodes 93 linguistic information from large corpora into vector representations or word embeddings. 94 These have been proven powerful in language modeling and some studies have suggested to 95 use them for AD detection due to excellent performance in a binary AD vs. control comparison 96 based on the ADReSS dataset (Balagopalan et al., 2021; Jawahar et al., 2019). Roshanzamir et 97 al. (2021) obtained an accuracy of 88.08% with BERT as an encoder with a logistic regression classifier, in a classification of AD vs. controls with English data from the Pitt corpus. Using 98 99 Hungarian data, Gosztolya et al. (2019) achieved 74%–82% accuracy in classifying diagnostic groups (cognitively healthy, MCI, and AD) based on speech (acoustic) features, and similar 100 101 results using language features.

Despite a number of promising studies, several challenges need to be addressed before speech- and language-based classification can be utilized in clinical applications. First, existing studies have mainly used the dataset from the InterSpeech challenge and its source corpus (Pitt Corpus) (Luz et al., 2020, 2021). These datasets only comprise English data from control and AD groups, so lacks data from disease stages in between, especially prodromal ones. Validation of the technically most advanced classifiers across different languages and the entire AD continuum is necessary. Second, current studies have investigated different linguistic levels with generic linguistic variables in a bottom-top fashion (Chapin et al., 2022), yet have not assessed in detail the differential feature performance across domains. To the best of our knowledge, only some studies have compared features from both speech and text, and they have not engaged in fine-grained comparisons of domains within these modalities, especially for the textual one, which has been represented chiefly by pretrained models (Balagopalan et al., 2020; Cummins et al., 2020; Zhu et al., 2021).

115 Finally, most available studies have elicited connected speech through a picture description, 116 using the Cookie Theft picture (Goodglass et al., 1972). This simple but efficient task has 117 significantly contributed to insights on AD detection through connected speech, but received 118 some criticism including the invocation of stereotypes of family life, elicitation of overly 119 simplified speech, and limited recollection and engagement (Berube et al., 2019; Clarke et al., 120 2021; Sherratt & Bryan, 2019). Beyond these, the Cookie Theft picture does not challenge the 121 creative and imaginative use of language as speech is generated while looking at the picture and objects are visually available for naming. As a hallmark of language is referencing objects 122 123 that are not visually present, tasks without visual prompts may show more sensitivity than 124 picture descriptions by challenging language production in one of its core features, namely 125 displaced reference. In particular, scene constructions (SC) have been proposed to stimulate 126 speech for richer information and better discrimination (Irish et al., 2015). SC requires a mental time travel to another location (e.g., a tropical island) and the imaginative representation of 127 directly experienced events at this location. As this information has an episodic and first-128 129 personal character, the task taps into a cognitive process widely reported to be impaired in AD

130 (Hassabis & Maguire, 2007; Schacter et al., 2017; Thakral et al., 2020).

In the present study we aimed (i) to test for the generalizability of previous results of automatic classification of AD using machine learning from English to Spanish/Catalan in a new dataset; (ii) to assess the performance of the classifier across a comprehensive number of language-related featural domains, and (iii) to include different MCI and preclinical groups at risk of AD in the classification.

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137 Methods

138 Dataset

We recruited 119 participants at the Memory Clinic from Ace Alzheimer Center Barcelona. 139 140 All of them were native Spanish and/or Catalan speakers. The referral center ethics committee 141 (Hospital Clínic i Provincial de Barcelona) approved the patient recruitment. Collection 142 protocols were under ethical standards according to the World Medical Association Declaration 143 of Helsinki - Ethical Principles for Medical Research Involving Human Subjects. Participants were diagnosed as cognitively healthy older controls (HOC), SCD, non-amnestic MCI 144 145 (naMCI), amnestic MCI (aMCI), and probable AD dementia (pAD). Briefly, SCD refers to the self-perception of cognitive problems, including memory loss, without impairment on the 146 147 standardized cognitive test (Jessen et al., 2014), while MCI implies that one or more cognitive 148 domains are impaired on the standardized cognitive test but activities of daily living are 149 preserved, according to Petersen's criteria (Petersen, 2004). Most SCD participants were part 150 of the FACEHBI study (Rodriguez-Gomez et al., 2017). Supplementary Information-A (SI-A, 151 similar below) specifies details about recruitment, diagnostic criteria, neuropsychological

assessment, and other issues. Table 1 shows the demographic and neuropsychological data ofthe sample (data: median (interquartile range, IQR)).

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155 Speech data elicitation and processing

Speech data were elicited through a scene construction task adapted from previous studies (Hassabis et al., 2007; Irish et al., 2015). Participants were instructed to construct a scene they imagined to witness and describe it with as much detail as possible, choosing one prompt from three options: "You are lying on the beach in a tropical bay."; "You are in a house that has been abandoned for many years."; and "You are in a circus tent". Most participants chose the first prompt. Speech samples were cut off after three minutes and transcribed into texts manually by a single researcher. The instructions for interview transcripts are shown in SI-B.

163 We extracted linguistic features from multiple domains, four directly from the audios and three from the transcripts (see Table 2). An overarching aim in feature selection was 164 165 comprehensiveness, in the sense that we wanted to comparatively assess all major levels of 166 organization of language/speech, including automatically extractable (i) acoustic spectral 167 coefficients, (ii) acoustic cepstral coefficients, (iii) voice quality, (iv) prosodic and (v) morpholexical features (the latter at the interface of the lexicon and morpho-syntax), (vi) manually 168 169 extractable syntactic features, and finally (vii) semantic ones (insofar as we can approximate 170 the latter with current NLP-technologies). In all of the cases (i) to (vii), there is some evidence 171 from previous literature to assume that the features might be discriminative.

In particular, we used openSMILE 3.0 to extract spectral, cepstral, voice quality, and part
of prosodic features from the CompareE 2016 feature set at the level of functionals (Eyben et

174 al., 2010; Schuller et al., 2016). Spectral and cepstral coefficients reveal the mathematical 175 properties of the sound waves, while voice quality features involve both phonatory and 176 resonatory characteristics. These three features process speech as sound waves and have been widely used in previous studies, demonstrating powerful detection capacities and high 177 178 robustness (Haider et al., 2020; Nasrolahzadeh et al., 2016; Thomas et al., 2020). Prosodic 179 features reflect suprasegmental patterns in how a speaker combines individual sounds into 180 integrated sequences with stress and intonation. Speech prosody has proved to be a sensitive measure to cognitive decline for AD even at early onset (Lofgren & Hinzen, 2022; Pistono et 181 182 al., 2016) and can contribute to accurate automated classification (Themistocleous et al., 2018). 183 As the prosodic profile in CompareE 2016 is not complete, we used Prosogram 3.0.1 to extract 184 more prosodic information, such as pitch variation and pitch stylization (Mertens, 2004). These 185 acoustic-prosodic measures are extracted in a fully automated form, not requiring human 186 transcription and annotation, and they are time-efficient (less than one minute to get all features 187 for each audio) manner, giving them special practical significance.

Syntactic features were manually annotated following the method of Chapin et al. (2022), 188 189 which targeted specific forms of syntactic complexity involved in referencing objects and 190 events. This feature set, though showing a close relation to neurodegeneration in AD, is not 191 automatable yet. Thus, we added the morpho-lexical features, which have often been used to 192 approximate syntax for measuring language changes in AD, especially in the context of NLP 193 (verb inflection in Fyndanis et al., 2011; ratios of different word classes in Guinn et al., 2014; 194 verb aspect in Manouilidou et al., 2020; verb voice in Nasiri et al., 2022). Morpho-lexical 195 features include the ratios of different word classes and the morphological variants of the

196 content words, which are automatically extracted by the Spanish and Catalan models from 197 Stanza 1.3.0. Lexical and morphological features were found to be important in the progression 198 of AD, in both contexts of group comparison (Kavé & Levy, 2003) and machine learning 199 (Evigoz et al., 2020). In addition, morphological variants in inflectional languages like Spanish and Catalan provide paths to observe grammar at word level (e.g. aspect and modality). 200 201 Semantic changes are thought to be prominent in AD, and large computational language 202 models capture distributional aspects of language as a proxy of meaning. A number of studies has found the application of language models on AD detection to be satisfying (Agbavor & 203 204 Liang, 2022; Balagopalan et al., 2021). The robust optimized version of BERT (RoBERTa) 205 was chosen for encoding the human transcripts. Due to the limitation on token numbers, we truncated the text and only encoded the first 510 tokens, if the text length was longer than that 206 207 (Liu et al., 2019). We used Catalan BERTa (RoBERTa-base) for Catalan data (https://huggingface.co/projecte-aina/roberta-base-ca-cased-tc) and Byte-Pair 208 Encoding 209 RoBERTa for Spanish data (https://huggingface.co/PlanTL-GOB-ES/roberta-base-bn) 210 (Armengol-Estapé et al., 2021; Gutiérrez-Fandiño et al., 2022). As RoBERTa models return an 211 embedding for every token in the text, which does not fit with the purpose of assigning a single 212 label to the text, we utilized the pooled output as a semantic representation. This output is the 213 embedding of the initial classification token that arises from the sequence output with 214 contextual information from all tokens of the sequence embedded in it. It is standardly used for 215 classification tasks. The complete list of feature sets is available in SI-C. In addition, we also 216 concatenated all the above-mentioned features into a long array as a comprehensive 217 representation of all features together.

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219 Experimental setup

220 We carried out several classifications involving different comparison groups, moving from 221 broader divisions to more fine-grained comparisons on the AD continuum: ten binary and one 222 ternary classification tasks. The binary comparisons were, firstly, the combined preclinical 223 group (HOC+SCD = CON) vs. the clinical groups (MCI and pAD separately), and the general 224 pathology (PATH) group comprising MCI and pAD together. Next, we performed three further 225 binary classifications: HOC from pAD, SCD from pAD, and HOC from SCD; and four 226 involving the MCI group: MCI from pAD, aMCI from pAD, naMCI from pAD, and aMCI 227 from naMCI. The ternary classification attempted to discriminate CON, MCI, and pAD. 228 Classification experiments were completed with the scikit-learn 1.0.2.

229 The random forests algorithm served as the classifier. This is an ensemble learning method 230 for constructing multiple decision trees to vote for the final label. Its robustness to overfitting 231 and noises motivated our choice (Breiman, 2001). Instead of forwarding all features to the 232 classifier, we selected only the most informative variables to reduce computational load, lower 233 the risk of overfitting, and remove noises from the feature set. In each experiment, we thus computed the ANOVA F-value between features in the feature set and ordered the features 234 235 based on these F values. The number of selected features was automatically determined based 236 on the classifier's performance, with a maximum of 1500 features. The classifier was evaluated 237 using ten-fold cross-validation with precision, recall, F1, and accuracy scores averaged across 238 the ten folds. Considering the imbalance between the number of participants in each 239 comparison group, we averaged the performance scores across comparison groups (i.e., the 240 macro scores). The macro F1 scores served as the major indicator for classifier performance as

241 it takes data distribution into account and compensates for group imbalance.

242

243 *Statistics*

244 To test the power of the classifier in distinguishing between different stages of AD, a repeated 245 measure ANOVA (RMANOVA) across different group comparisons was carried out. In 246 addition, we conducted a RMANOVA across the different speech and language domains, to investigate how different feature sets influence the performance of the machine learning 247 248 classifier. For each RMANOVA, we tested the assumption of sphericity and corrected with 249 Greenhouse-Geisser method when the assumption was violated. Post-hoc tests were conducted 250 in case of significant difference with Holm adjusted p-values. All statistical results are rounded 251 to three digits, SD stands for standard deviation. Furthermore, we categorized feature sets into two overall modalities with the equal number of features, one 'speech' modality including the 252 253 four features directly extracted from audios, and one 'text'-modality including the three feature sets from transcripts and the concatenation of all features. We carried out a paired-sample 254 255 Wilcoxon signed-rank test to check if the speech modality and textual modality are different from each other. Analyses were run with JASP 0.16.3.0. 256

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258 **Results**

259 Classification performance scores across groups

260 Table 3 reports the F1 scores of the random forest classifier on the ten classification tasks. To

261 make the table more concise, we report only the averaged F1 scores performance score across

262 groups for classification. The complete group-wise performance matrices, including precision, 263 recall, F1, and accuracy scores, are reported in SM-D. Figure 1 (a) shows the distribution of 264 the F1 scores obtained by the classifier on different group comparisons using different 265 linguistic feature sets, and (b) shows the violin plot of F1 scores for each comparison, with a 266 line of mean F1 scores. The post-hoc comparisons for both groups and linguistic domains are 267 shown in SM-E. As indicated by the RMANOVA, the classifier performed significantly different among different comparisons (F(10) = 14.423, p < .001, $\eta^2 = .673$). Measured with 268 269 the average F1 scores, compared to the ternary classification (mean = .572, SD = .077), the classifier performed significantly better on all binary classifications (p < .05). For binary 270 271 classifications, the classifier performed best on distinguishing between groups without (HOC and SCD) and with (MCI and pAD) cognitive impairment, specifically SCD from pAD (mean 272 273 = .878, SD = .034), followed by CON (that is, the combined group without objective cognitive decline) from pAD (mean = .812, SD = .039), CON from the combined 'pathological' group 274 275 with cognitive decline (PATH) (mean = .786, SD = .024), and CON from MCI (mean = .774, 276 SD = .038). Performance on the three comparisons between CON and pathological groups 277 (pAD and MCI) were similar to each other (p = 1.000). Next, the classifier distinguished HOC from pAD (mean = .758, SD = .053) and HOC from SCD (mean = .749, SD = .143). It is 278 279 noteworthy that performance on SCD vs. pAD was significantly better than HOC vs. pAD (p 280 = .005 < .05). The seventh to tenth performance scores were: aMCI vs. naMCI (mean = .747, 281 SD = .101), naMCI vs. pAD (mean = .720, SD = .068), MCI vs. pAD (mean = .695, SD = .061), 282 and finally aMCI vs. pAD (mean = .678, SD = .158). The performance on naMCI vs. pAD was 283 almost the same as that of MCI vs. pAD (p = 1.000) and aMCI vs. pAD (p = 1.000). The

284 performance on CON vs. MCI was not significantly better than these four comparisons among 285 MCI groups and between them and pAD (p > .05).

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287 Classification performance scores across linguistic levels and modalities

288 Figure 1 (c) shows the violin plot of F1 scores for each feature set with a line of mean F1 scores. 289 As indicated by the RMANOVA with Greenhouse-Geisser sphericity correction, the machine 290 learning classifier performed significantly different among different feature sets (F(2.214) =12.423, p < .001, $\eta^2 = .554$). Ordered by mean F1 scores, the classifier performed best on the 291 concatenation of all features (mean = .813, SD = .069), followed by the voice quality 292 293 measurements (mean = .796, SD = .075), the spectral coefficients (mean = .791, SD = .078), 294 the cepstral coefficients (mean = .773, SD = .087), the embeddings from RoBERTa (mean 295 = .738, SD = .111), the prosodic features (mean = .724, SD = .097), the morpho-lexical features 296 (mean = .659, SD = .104), and the syntactic features (mean = .648, SD = .131). Syntactic and 297 morpho-lexical features were similar to each other (p = 1.000), and significantly worse than all 298 other feature sets (p < .05) except prosodic features (p = .058, p = .158, respectively). Prosodic 299 features performed significantly worse only relative to the concatenation of all features (p = .013, p < .05). Post-hoc comparisons among other pairs of feature sets were all insignificant 300 301 (p > .05). After grouping linguistic levels into two modalities, speech and text, speech-based 302 features (mean = .771, median = .776, SD = .087) discriminated groups significantly better than text-based ones (mean = .714, median = .736, SD = .123, z = 2.019, p = .043 < .05). Figure 303 304 1 (d) shows the violin plot of F1-scores for each modality with a line of mean F1 scores.

306 Discussion

307 This study aimed to (i) test for the generalizability of previous results of automatic classification of AD from English to Spanish/Catalan in a new dataset; (ii) assess the 308 309 performance of the classifier across different language-related featural domains, and (iii) 310 include different MCI as well as preclinical groups at risk of AD in the classification. Our 311 results confirm that similar performance as in previous studies based on Hungarian and English 312 data can be generalized on our new Spanish/Catalan data (Gosztolya et al., 2019; Haulcy & 313 Glass, 2021). For all binary comparisons we made, most of the F1 scores are around or higher than .7. For some specific separations we even achieved more impressive results, such as the 314 315 F1 score of .912 in separating HOC from pAD based on spectral coefficients. For ternary 316 classification, though the performance is worse, between .447-.662 by F1 score, they were 317 higher than the chance level of .330, with the highest scores doubling the latter. These results 318 suggest that speech analysis can be a potentially powerful and generalizable approach for 319 automated pAD detection and risk for it.

320 As for performance in different feature domains, accuracies were generally satisfactorily 321 high across domains, with the exception of morpho-lexical features and syntactic features. The former finding may be expected, as declarative memory-related cognitive impairment in AD 322 323 may not result in changes in morphology, associated with procedural memory. More surprising 324 is the finding on syntactic features, in light of the study of Chapin et al. (2022), where a number 325 of hand-selected syntactic measures related to hierarchical syntactic complexity discriminated 326 between controls, MCI and AD groups. One possibility is that significant changes in syntactic impairment occur in MCI, as the F1 scores achieved on distinguishing controls from MCIs and 327

328 pAD were all around or above .7 (.688-.860), but less than .6 when comparing within groups 329 with or without (objective) cognitive impairment (.460-.589). Decline in syntactic complexity 330 also seems to be a late effect in the pathophysiological process, as compared to speech domains. 331 Thus, in all three textual domains investigated, RoBERTa performed significantly better than syntactic and morpho-lexical features, and comparably to speech domains. Unlike manually 332 333 designed feature sets including our syntactic one, RoBERTa is an integrated linguistic 334 representation wrapping up what the model learned from the pre-trained corpus and current contexts into the embeddings. This highlights the importance of semantic changes in AD, as 335 336 RoBERTa originates from distributional semantics-based word embeddings. Nonetheless, 337 current studies have shown that these BERT-based models also capture lexical, syntactic, and 338 conversational information in addition to semantic information (Kumar et al., 2021; Staliūnaitė 339 & Iacobacci, 2020). Syntactic changes at the phrasal level could be important in AD, even for 340 the early stages. Again, morphology and phrasal syntactic complexity could be more a matter 341 of procedural memory, while the syntactic variables in our syntactic feature set could be argued 342 to capture more declarative aspects of language use, such as specific forms of complexity 343 needed to express episodic semantic information.

Another unexpected finding was that performance in speech domains was significantly better than in textual domains. This raises the thought-provoking question whether we even need transcripts and textual analysis, given the impressive performances of acoustic features and the high costs of the transcription task. Although fusing speech and text domains gives slight increases (less than .1) in performance scores, it is questionable whether such small increases balance the cost of transcription. The voice quality measures and spectral and cepstral 350 coefficients performed the best and similar to each other. Although previous studies 351 hypothesized voice quality changes as an ex-post effect from the physiological impairment of 352 the fine control and the slowing down of vocal organs due to MCI and AD, the F1 scores of .742 353 on separating HOC and SCD and .814 on separating aMCI and naMCI suggest the roles of 354 cognitive decline and memory loss in the changes of voice quality (Themistocleous et al., 2020). 355 Paralinguistic features have been verified as performative in AD classification for multiple 356 languages (Lindsay et al., 2021) and even across languages (Martinez de Lizarduy et al., 2017), which has been confirmed in our study. Similar to the voice quality measures, some studies 357 358 also related paralinguistic features to voicing handicaps, so treated these changes as a side 359 effect of AD (Awan et al., 2014). However, we found that the spectral and cepstral coefficients 360 were more discriminative when separating HOC and SCD, followed by separations between 361 individuals with and without cognitive impairment, including the aMCI vs naMCI, and finally 362 between MCI and pAD. A more reasonable interpretation could be that paralinguistic features 363 represent variance from other factors such as affective, apathy and executive functions (Lindsay et al., 2021). 364

As for prosody, when ranking the performance of prosody and syntax on different classification tasks from highest to lowest, similar results were found between them, with the comparisons between groups with and without cognitive impairment at the top and comparisons within these two general groups at the bottom. As prosody and syntax respectively represent the unification of sounds and words, we may conclude that organization abilities in these two domains decline greatly from cognitively healthy to cognitive impairment, but slowly progress within these two general phases. 372 Our final aim was to test the extendibility of previous classification results to further 373 groups on the AD spectrum, specifically SCD and different MCI groups. Remarkably, very 374 high accuracies were obtained when classifying cognitively healthy individuals with and 375 without cognitive complaints (HOC and SCD), and classifying each from pAD, specifically 376 when using the purely speech-based feature domains - spectral and cepstral coefficients. To 377 our knowledge, this is the first report of an automatized differentiation between these 378 preclinical groups, which is particularly noteworthy insofar as it does not depend on 379 transcription. Future work following people with SCD over time could investigate this issue, 380 by comparing the speech of converting vs. non-converting SCDs after an interval. Equally 381 striking in our results is the differentiability of the amnestic and non-amnestic MCI groups, 382 again based on spectral and cepstral features. Despite the high differentiability, although aMCI 383 and naMCI are not that similar between themselves, they are similarly different from AD, unlike SCD and HOC. Furthermore, similar patterns were observed in groups, linguistic 384 385 domains, and linguistic modalities, when we applied the gradient boosting algorithm as the 386 classifier, another ensemble learning algorithm where the decision trees are not independent 387 but will correct each other. Results from this algorithm and statistical comparisons can be found in SI-F. These similar patterns from a different algorithm indicate robustness in our 388 389 classification results.

This study has several limitations. First, the dataset is relatively small from a computational perspective, so we can neither train the model with large data nor use state-ofthe-art deep learning techniques. Although we managed to validate the result by using crossvalidation, this still limits the performance of the classifier. Secondly, it is impossible to capture 394 the full picture in every linguistic domain, though we used the available and already verified 395 features to ensure representativeness. Finally, the performance on ternary classification is not 396 high, likely due to the heterogeneity in groups and the size of the dataset.

397 In conclusion, our study shows that using machine learning based on speech can be a 398 potentially powerful tool for detecting cognitive impairment and its gradation based on pAD 399 pathology and that different linguistic domains play significantly different roles in this 400 procedure. With clinical applications in mind, we underline both the high performance of speech-based measures as compared to text-based ones, and the discriminability even of 401 402 objectively unimpaired healthy older adults with and without SCD, and of groups with 403 amnestic and non-amnestic MCI. Combined with other behavioral markers or biological ones 404 such as blood or retinal appearances, speech analysis may well prove to provide essential help 405 for establishing early and robust diagnostic markers of AD, which are inexpensive and widely available. 406

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434 Data Availability Statement

The datasets generated and analyzed during the current study are not publicly available due to
ethical requirements. Codes for processing the data and generating the figures are available
from the corresponding author upon request.

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665 Supplementary file description:

The supplementary information is divided into six parts : (A) The recruitment procedure,
diagnostic criteria, neuropsychological battery, impact of the epidemic, and criteria for
inclusion and exclusion. (B) Speech transcription instruction. (C) Feature list and definitions.
(D) Detailed group-wise classifier performance from the random forest. (F) Post-hoc analysis
of the RMANOVA tests. (F) Results and statistical comparison from Gradient Boosting.

672 Tables

	HOC	SCD	naMCI	aMCI	pAD	Test	<i>p</i> value
Number	18	31	23	16	31	/	/
Age	66 (8)	69 (10)	75 (8)	78 (7)	81 (9)	KW test	<.001***
Age Range	58 - 89	56 - 85	52 - 88	66 - 89	60 - 93	/	/
Sex	61.1	25.8	39.1	37.5	45.2	χ^2	.177
Education	7 (4)	7 (3)	5 (3)	5 (3)	5 (2)	KW test	.000***
Language	33.3	22.6	4.4	12.5	12.9	χ^2 (Fisher)	.126
	29.0	29.0	27.0	28.0	23.0	VW toot	< 001***
WIMSE score	(1.0)	(2.0)	(3.0)	(2.1)	(4.5)	K w test	< .001****
CDR score	0	0	.5	.5	1 or 2	/	/

673 **Table 1**: Participant demographics[†]

Note: HOC: healthy older control; SCD: subjective cognitive decline; naMCI: non-amnestic
mild cognitive impairment; aMCI: amnestic mild cognitive impairment; pAD: probable
Alzheimer's disease dementia; KW test: Kruskal-Wallis test; MMSE: Mini-Mental State
Examination; CDR: Clinical Dementia Rating.

678 *p < .05, **p < .01, and ***p < .001.

679 †: Age in years. Gender is represented by the proportion of females. Education in education
680 level. Language is represented by the proportion of people answering questions in Catalan.
681 Fisher's exact test is applied for language as several expected frequency less than 5.

Levels	Description	Num	Tools
Spectral	Auditory spectrum and the relative spectral	2800	openSMILE
	transform		
Cepstral	Mel-Frequency cepstral coefficients 0-14.	1400	openSMILE
Voice quality	Jitter, shimmer, loudness, and log harmony-	2012	openSMILE
	noise-ratio		
Prosodic	Include frequency, speech rate, pitch	199	openSMILE&
	variation, pitch stylization etc.		Prosogram
Morpho-	Ratios of different word classes and the	154	Stanza
lexical	morphological variants, e.g. masculine nouns		
Syntactic	Ratios of syntactic features selected from	21	Manual
	Chapin et al.(Chapin et al., 2022), e.g. verb		
	modality		
Semantic	Pooled output of RoBERTa models	768^{\dagger}	RoBERTa

Table 2: Linguistic features extracted

684 †: number of dimensions of the word embeddings

Comparison	spectral	cepstral	prosodic	voice quality	syntactic	morpho-lexical	RoBERTa	all
CON/PATH	.789	.776	.805*	.804*	.802*	.738	.768	.806*
CON/pAD	.825*	.833*	.819*	.853**	.771	.735	.826*	.834*
CON/MCI	.801*	.769	.798	.799	.743	.700	.771	.814*
HOC/pAD	.765	.775	.757	.800*	.688	.676	.832*	.771
HOC/SCD	.912***	.908***	.702	.742	.534	.649	.649	.900***
HOC/pAD	.765	.775	.757	.800*	.688	.676	.832*	.771
SCD/pAD	.878**	.880**	.854**	.903***	.860**	.819*	.928***	.898**
MCI/pAD	.732	.724	.682	.753	.589	.635	.675	.768
aMCI/pAD	.719	.724	.539	.866**	.460	.508	.737	.869**
naMCI/ pAD	.734	.764	.718	.801*	.581	.690	.696	.776
aMCI/ naMCI	.881**	.776	.696	.814*	.581	.656	.730	.847*

Table 3: Classifier performance across different comparisons and feature sets

CON/MCI/pAD	.662	.579	.591	.618	.516	.447	.505	.660
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686 Note: *** F1 score > = .9 for very good, ** F1 score > = .85 for good, * F1 score > = .8 for not bad

688 Figures

- 689 **Figure 1.** (a) Scatter plot of F1-scores across different feature sets and classification tasks; (b)
- 690 Means, standard deviations and distributions of the same F1-scores across classification tasks
- 691 with the mean line; (c) F1-scores across different feature sets with the mean line; (d) F1-scores
- 692 between different modalities (speech vs. text) with the mean line.
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694	Learning	Outcomes
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695	•	Machine learning based on automatically extracted language features detected cognitive
696		decline from early stages of the AD continuum in a new Spanish-Catalan dataset.

Different speech and language domains showed differential discrimination performance
 between groups, with features extracted directly from speech performing better than those
 from the text.

• Before the onset of objective cognitive impairment, speech and language from older adults

701 with Subjective Cognitive Decline (SCD) showed speech and language differences from

702 controls without SCD, indicating potential heterogeneity in these non-clinical groups.