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Benchmarking of eight recurrent neural network variants for breath phase and adventitious sound detection on a self-developed open-access lung sound database—HF_Lung_V1 --Manuscript Draft--

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Abstract:	<p>A reliable, remote, and continuous real-time respiratory sound monitor with automated respiratory sound analysis ability is urgently required in many clinical scenarios—such as in monitoring disease progression of coronavirus disease 2019—to replace conventional auscultation with a handheld stethoscope. However, a robust computerized respiratory sound analysis algorithm has not yet been validated in practical applications. In this study, we developed a lung sound database (HF_Lung_V1) comprising 9,765 audio files of lung sounds (duration of 15 s each), 34,095 inhalation labels, 18,349 exhalation labels, 13,883 continuous adventitious sound (CAS) labels (comprising 8,457 wheeze labels, 686 stridor labels, and 4,740 rhonchi labels), and 15,606 discontinuous adventitious sound labels (all crackles). We conducted benchmark tests for long short-term memory (LSTM), gated recurrent unit (GRU), bidirectional LSTM (BiLSTM), bidirectional GRU (BiGRU), convolutional neural network (CNN)-LSTM, CNN-GRU, CNN-BiLSTM, and CNN-BiGRU models for breath phase detection and adventitious sound detection. We also conducted a performance comparison between the LSTM-based and GRU-based models, between unidirectional and bidirectional models, and between models with and without a CNN. The results revealed that these models exhibited adequate performance in lung sound analysis. The GRU-based models outperformed, in terms of F1 scores and areas under the receiver operating characteristic curves, the LSTM-based models in most of the defined tasks. Furthermore, all bidirectional models outperformed their unidirectional counterparts. Finally, the addition of a CNN improved the accuracy of lung sound analysis, especially in the CAS detection tasks.</p>
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1 **Benchmarking of eight recurrent neural network variants for breath phase and**
2 **adventitious sound detection on a self-developed open-access lung sound**
3 **database—HF_Lung_V1**

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24

25 **ABSTRACT**

26 A reliable, remote, and continuous real-time respiratory sound monitor with automated respiratory
27 sound analysis ability is urgently required in many clinical scenarios—such as in monitoring disease
28 progression of coronavirus disease 2019—to replace conventional auscultation with a handheld
29 stethoscope. However, a robust computerized respiratory sound analysis algorithm has not yet been
30 validated in practical applications. In this study, we developed a lung sound database (HF_Lung_V1)
31 comprising 9,765 audio files of lung sounds (duration of 15 s each), 34,095 inhalation labels, 18,349
32 exhalation labels, 13,883 continuous adventitious sound (CAS) labels (comprising 8,457 wheeze
33 labels, 686 stridor labels, and 4,740 rhonchi labels), and 15,606 discontinuous adventitious sound
34 labels (all crackles). We conducted benchmark tests for long short-term memory (LSTM), gated
35 recurrent unit (GRU), bidirectional LSTM (BiLSTM), bidirectional GRU (BiGRU), convolutional
36 neural network (CNN)-LSTM, CNN-GRU, CNN-BiLSTM, and CNN-BiGRU models for breath
37 phase detection and adventitious sound detection. We also conducted a performance comparison
38 between the LSTM-based and GRU-based models, between unidirectional and bidirectional models,
39 and between models with and without a CNN. The results revealed that these models exhibited
40 adequate performance in lung sound analysis. The GRU-based models outperformed, in terms of *F1*
41 scores and areas under the receiver operating characteristic curves, the LSTM-based models in most of
42 the defined tasks. Furthermore, all bidirectional models outperformed their unidirectional counterparts.
43 Finally, the addition of a CNN improved the accuracy of lung sound analysis, especially in the CAS

44 detection tasks.

45

46 *Keywords:* adventitious sound, auscultation, convolutional neural networks, lung sound, recurrent

47 neural networks, respiratory monitor

48 **1. Introduction**

49 Respiration is vital for the normal functioning of the human body. Therefore, clinical physicians
50 are frequently required to examine respiratory conditions. Respiratory auscultation [1-3] using a
51 stethoscope has long been a crucial first-line physical examination. The chestpiece of a stethoscope
52 is usually placed on a patient's chest or back for lung sound auscultation or over the patient's
53 tracheal region for tracheal sound auscultation. During auscultation, breath cycles can be inferred,
54 which help clinical physicians evaluate the patient's respiratory rate. In addition, pulmonary
55 pathologies are suspected when the frequency or intensity of respiratory sounds changes or when
56 adventitious sounds, including continuous adventitious sounds (CASs) and discontinuous
57 adventitious sounds (DASs), are identified [1, 2, 4]. Patients with coronavirus disease 2019 exhibit
58 adventitious sounds [5]; hence, auscultation may be a useful approach for disease diagnosis [6] and
59 disease progression tracking. However, auscultation performed using a conventional handheld
60 stethoscope involves some limitations [7]. First, the interpretation of auscultation results
61 substantially depends on the subjectivity of the practitioners. Even experienced clinicians might not
62 have high consensus rates in their interpretations of auscultatory manifestations [8, 9]. Second,
63 auscultation is a qualitative analysis method. Comparing auscultation results between individuals and
64 quantifying the sound change by reviewing historical records are difficult tasks. Third, prolonged
65 continuous monitoring of respiratory sound is almost impractical.

66 To overcome the aforementioned limitations, computerized methods for respiratory sound
67 recording and analyses based on traditional signal processing and machine learning have been
68 proposed and reviewed [4, 10-13]. With the advent of the deep learning era, studies have developed
69 novel deep learning-based methods for respiratory sound analysis. However, many of such studies
70 have focused on only distinguishing healthy participants from participants with respiratory disorders
71 [14-18] and distinguishing various types of normal breathing sounds from adventitious sounds
72 [19-25]. Only a few studies [26-29] have explored the use of deep learning for detecting breath
73 phases and adventitious sounds. Moreover, most previous studies on computerized lung sound
74 analysis have been limited by insufficient data. As of writing this paper, the largest reported
75 respiratory sound database is ICBHI 2017 Challenge [30], which comprises 6,898 breath cycles and
76 10,775 events of wheezes and crackles acquired from 126 individuals.

77 Data size plays a major role in the creation of a robust and accurate deep learning-based respiratory
78 sound analysis algorithm [31, 32]. Accordingly, the first aim of the present study was to establish a
79 large and open-access respiratory sound database for training such algorithms for the detection of
80 breath phase and adventitious sounds, mainly focusing on lung sounds. The second aim was to conduct
81 a benchmark test on the established lung sound database by using eight recurrent neural network
82 (RNN)-based models. RNNs [33] are effective for time-series analysis; long short-term memory
83 (LSTM) [34] and gated recurrent unit (GRU) [35] networks, which are two RNN variants, exhibit
84 superior performance to the original RNN model. However, whether LSTM models are superior to

85 GRU models (and vice versa) in many applications, particularly in respiratory sound analysis, is
86 inconclusive. Bidirectional RNN models [36, 37] can transfer not only past information to the future
87 but also future information to the past; these models consistently exhibit superior performance to
88 unidirectional RNN models in many applications [38-40] as well as in breath phase and crackle
89 detection [29]. However, whether bidirectional RNN models outperform unidirectional RNN models in
90 CAS detection has yet to be determined. Furthermore, the convolutional neural network (CNN)–RNN
91 structure has been proven to be suitable for heart sound analysis [41], lung sound analysis [19], and
92 other tasks [39, 42]. Nevertheless, the application of the CNN–RNN structure in respiratory sound
93 detection has yet to be fully investigated. Benchmarking can enable demonstrating the reliability and
94 goodness of a database; it can also be applied to investigate the performance of the RNN variants in
95 respiratory analysis.

96 In summary, the aims of this study are outlined as follows:

- 97 ■ Establish the largest open-access lung sound database as of writing this paper—HF_Lung_V1
98 (https://gitlab.com/techsupportHF/HF_Lung_V1).
- 99 ■ Conduct a performance comparison between LSTM and GRU models, between unidirectional and
100 bidirectional models, and between models with and without a CNN in breath phase and
101 adventitious sound detection based on lung sound data.
- 102 ■ Discuss factors influencing model performance.

103

104 **2 Establishment of the lung sound database**

105 *2.1 Data sources and patients*

106 The lung sound database was established using two sources. The first source was a database
107 used in a datathon in Taiwan Smart Emergency and Critical Care (TSECC), 2020, under the license
108 of Creative Commons Attribution 4.0 (CC BY 4.0), provided by the Taiwan Society of Emergency
109 and Critical Care Medicine. Lung sound recordings in the TSECC database were acquired from 261
110 patients.

111 The second source was sound recordings acquired from 18 residents of a respiratory care ward
112 (RCW) or a respiratory care center (RCC) in Northern Taiwan between August 2018 and October
113 2019. The recordings were approved by the Research Ethics Review Committee of Far Eastern
114 Memorial Hospital (case number: 107052-F). Written informed consent was obtained from the 18
115 patients. This study was conducted in accordance with the 1964 Helsinki Declaration and its later
116 amendments or comparable ethical standards.

117 All patients were Taiwanese and aged older than 20 years. Descriptive statistics regarding the
118 patients' demographic data, major diagnosis, and comorbidities are presented in Table 1; however,
119 information on the patients in the TSECC database is missing. Moreover, all 18 RCW/RCC residents
120 were under mechanical ventilation.

121

122 **Table 1. Demographic data of patients.**

	Subjects from RCW/RCC	Subjects in TSECC Database
Number (n)	18	261
Gender (M/F)	11/7	NA
Age	67.5 (36.7, 98.3)	NA
Height (cm)	163.6 (147.2, 180.0)	NA
Weight (kg)	62.1 (38.2, 86.1)	NA
BMI (kg/m ²)	23.1 (15.6, 30.7)	NA
Respiratory Diseases		
ARF	4 (22.2%)	NA
CRF	8 (44.4%)	NA
COPD AE	1 (5.6%)	NA
COPD	2 (11.1%)	NA
Pneumonia	4 (22.2%)	NA
ARDS	1 (5.6%)	NA
Emphysema	1 (5.6%)	NA
Comorbidity		
CKD	1 (5.6%)	NA
AKI	3 (16.7%)	NA
CHF	2 (11.1%)	NA
DM	7 (38.9%)	NA
HTN	6 (33.3%)	NA
Malignancy	1 (5.6%)	NA
Arrythmia	1 (5.6%)	NA
CAD	1 (5.6%)	NA

123 RCW: respiratory care ward, RCC: respiratory care center, ARF: acute respiratory failure, CRF: chronic respiratory failure, COPD AE: chronic
124 obstructive pulmonary disease acute exacerbation, COPD: chronic obstructive pulmonary disease, ARDS: acute respiratory distress syndrome, CKD:
125 chronic kidney disease, AKI: acute kidney injury, CHF: chronic heart failure, DM: diabetes, HTN: hypertension, CAD: cardiovascular disease. The mean
126 values of the age, height, weight, and BMI are presented, with the corresponding 95% CI in parentheses.

127

128

129 2.2 Sound recording

130 Breathing lung sounds were recorded using two devices: (1) a commercial electronic

131 stethoscope (Littmann 3200, 3M, Saint Paul, Minnesota, USA) and (2) a customized multichannel
132 acoustic recording device (HF-Type-1) that supports the connection of eight electret microphones.
133 The signals collected by the HF-Type-1 device were transmitted to a tablet (Surface Pro 6, Microsoft,
134 Redmond, Washington, USA; Fig 1). Breathing lung sounds were collected at the eight locations
135 (denoted by L1–L8) indicated in Fig 2a. The auscultation locations are described in detail in the
136 caption of Fig 2. The two devices had a sampling rate of 4,000 Hz and a bit depth of 16 bits. The
137 audio files were recorded in the WAVE (.wav) format.

138

139

140 **Fig 1. Customized multichannel acoustic recording device (HF-Type-1) connected to a tablet.**

141

142 **Fig 2. Auscultation locations and lung sound recording protocol.** (a) Auscultation locations (L1–
143 L8): L1: second intercostal space (ICS) on the right midclavicular line (MCL); L2: fifth ICS on the
144 right MCL; L3: fourth ICS on the right midaxillary line (MAL); L4: tenth ICS on the right MAL; L5:
145 second ICS on the left MCL; L6: fifth ICS on the left MCL; L7: fourth ICS on the left MAL; and L8:
146 tenth ICS on the left MAL. (b) A standard round of breathing lung sound recording with Littmann
147 3200 and HF-Type-1 devices. The white arrows represent a continuous recording, and the small red
148 blocks represent 15-s recordings. When the Littmann 3200 device was used, 15.8-s signals were
149 recorded sequentially from L1 to L8. Subsequently, all recordings were truncated to 15 s. When the
150 HF-Type-1 device was used, sounds at L1, L2, L4, L5, L6, and L8 were recorded simultaneously.
151 Subsequently, each 2-min signal was truncated to generate new 15-s audio files.

152

153 All lung sounds in the TSECC database were collected using the Littmann 3200 device only,
154 where 15.8-s recordings were obtained sequentially from L1 to L8 (Fig 2b; Littmann 3200). One

155 round of recording with the Littmann 3200 device entails a recording of lung sounds from L1 to L8.

156 The TSECC database was composed of data obtained from one to three rounds of recording with the

157 Littmann 3200 device for each patient.

158 We recorded the lung sounds of the 18 RCW/RCC residents by using both the Littmann 3200

159 device and the HF-Type-1 device. The Littmann 3200 recording protocol was in accordance with that

160 used in the TSECC database, except that data from four to five rounds of lung sound recording were

161 collected instead. The HF-Type-1 device was used to record breath sounds at L1, L2, L4, L5, L6, and

162 L8. One round of recording with the HF-Type-1 device entails a synchronous and continuous

163 recording of breath sounds for 30 min (Fig 2b; HF-Type-1). However, the recording with the

164 HF-Type-1 device was occasionally interrupted; in this case, the recording duration was <30 min.

165 Voluntary deep breathing was not mandated during the recording of lung sounds. The statistics

166 of the recordings are listed in Table 2.

167

168 **Table 2. Statistics of recordings and labels of HF_Lung_V1 database.**

	Littmann 3200	HF-Type-1	Total
Subjects			
n	261	18	261
Recordings			
Filename prefix	steth_	trunc_	NA
Rounds of recording	748	70	NA
No of 15-sec recordings	4504	5261	9765
Total duration (min)	1126	1315.25	2441.25
Labels			
No of I	16535	17560	34095

Total duration of I (min)	257.17	271.02	528.19
Mean duration of I (sec)	0.93	0.93	0.93
No of E	9107	9242	18349
Total duration of E (min)	160.25	132.60	292.85
Mean duration of E (sec)	1.06	0.86	0.96
No of C/W/S/R	6984/3974/152/2858	6899/4483/534/1882	13883/8457/686/4740
Total duration of C/W/S/R (min)	105.90/63.92/1.94/40.04	85.26/55.80/7.52/21.94	191.16/119.73/9.46/61.98
Mean duration of C/W/S/R (sec)	0.91/0.97/0.76/0.84	0.74/0.75/0.85/0.70	0.83/0.85/0.83/0.78
No of D	7266	8340	15606
Total duration of D (min)	111.75	55.80	230.87
Mean duration of D (sec)	0.92	0.87	0.89

169 I: inhalation, E: exhalation, W: wheeze, S: stridor, R: rhonchus, C: continuous adventitious sound, D: discontinuous adventitious sound. W, S, and R were
170 combined to form C.

171
172

173 2.3 Audio file truncation

174 In this study, the standard duration of an audio signal used for inhalation, exhalation, and
175 adventitious sound detection was 15 s. This duration was selected because a 15-s signal contains at
176 least three complete breath cycles, which are adequate for a clinician to reach a clinical conclusion.
177 Furthermore, a 15-s breath sound was be used previously for verification and validation [43] .

178 Because each audio file generated by the Littmann 3200 device had a length of 15.8 s, we
179 cropped out the final 0.8-s signal from the files (Fig 2b; Littmann 3200). Moreover, we used only the
180 first 15 s of each 2-min signal of the audio files (Fig 2b; HF-Type-1) generated by the HF-Type-1
181 device. Table 2 presents the number of truncated 15-s recordings and the total duration.

182

183 2.4 Data labeling

184 Because the data in the TSECC database contains only classification labels indicating whether a
185 CAS or DAS exists in a recording, we attempted to label the event level of all sound recordings. Two
186 board-certified respiratory therapists (NJL and YLW) and one board-certified nurse (WLT), with 8, 3,
187 and 13 years of clinical experience, respectively, were recruited to label the start and end points of
188 inhalation (I), exhalation (E), wheeze (W), stridor (S), rhonchus (R), and DAS (D) events in the
189 recordings. They labeled the sound events by listening to the recorded breath sounds while
190 simultaneously observing the corresponding patterns on a spectrogram by using customized labeling
191 software [44]. The labelers were asked not to label sound events if they could not clearly identify the
192 corresponding sound or if an incomplete event at the beginning or end of an audio file caused
193 difficulty in identification. BFH held regular meetings to ensure that the labelers had good agreement
194 on labeling criteria based on a few samples by judging the mean pseudo- κ value [27]. When
195 developing artificial intelligence (AI) detection models, we combined the W, S, and R labels to form
196 CAS labels (C). Moreover, the D labels comprised only crackles, which were not differentiated into
197 coarse or fine crackles. The labelers were asked to label the period containing crackles but not a
198 single explosive sound (generally less than 25 ms) of a crackle. Each recording was annotated by
199 only one labeler; thus, the labels did not represent perfect ground truth. However, we used the labels
200 as ground-truth labels for model training, validation, and testing. The statistics of the labels are listed
201 in Table 2.

202

203 **3. Inhalation, exhalation, CAS, and DAS detection**

204 *3.1 Framework*

205 The inhalation, exhalation, CAS, and DAS detection framework developed in this study is
206 displayed in Fig 3. The prominent advantage of the research framework is its modular design.
207 Specifically, each unit of the framework can be tested separately, and the algorithms in different parts
208 of the framework can be modified to achieve optimal overall performance. Moreover, the output of
209 some blocks can be used for multiple purposes. For instance, the spectrogram generated by the
210 preprocessing block can be used as the input of a model or for visualization in the user interface for
211 real-time monitoring.

212

213 **Fig. 3. Pipeline of detection framework.**

214

215 The framework comprises three parts: preprocessing, deep learning–based modeling, and
216 postprocessing. The preprocessing part involves signal processing and feature engineering
217 techniques. The deep learning–based modeling part entails the use of a well-designed neural network
218 for obtaining a sequence of classification predictions rather than a single prediction. The
219 postprocessing part involves merging the segment prediction results and eliminating the burst event.

220

221 *3.2 Preprocessing*

222 We processed the lung sound recordings at a sampling frequency of 4 kHz. First, to eliminate
223 the 60-Hz electrical interference and a part of the heart sound noise, we applied a high-pass filter to
224 the recordings by setting a filter order of 10 and cut-off frequency of 80 Hz. The filtered signals were
225 then processed using the short-time Fourier transform (STFT). In the STFT, we set a Hanning
226 window size of 256 and hop length of 64; no additional zero-padding was applied. Thus, a 15-s
227 sound signal could be transformed into a corresponding spectrogram with a size of 938×129 . To
228 obtain the spectral information regarding the lung sounds, we extracted the following features [29,
229 45]:

- 230 ■ Spectrogram: We extracted 129-bin log-magnitude spectrograms.
- 231 ■ Mel frequency cepstral coefficients (MFCCs): We extracted 20 static coefficients, 20 delta
232 coefficients (Δ), and 20 acceleration coefficients (Δ^2). We used 40 mel bands within a frequency
233 range of 0–4,000 Hz. The frame width used to calculate the delta and acceleration coefficients
234 was set to 9, which resulted in a 60-bin vector per frame.
- 235 ■ Energy summation: We computed the energy summation of four frequency bands, namely 0–
236 250, 250–500, 500–1,000, and 0–2,000 Hz, and obtained four values per time frame.

237 After extracting the aforementioned features, we concatenated them to form a 938×193 feature
238 matrix. Subsequently, we conducted min–max normalization on each feature. The values of the
239 normalized features ranged between 0 and 1.

240

241 3.3 Deep learning models

242 We investigated the performance of eight RNN models, namely LSTM, GRU, bidirectional
243 LSTM (BiLSTM), bidirectional GRU (BiGRU), CNN-LSTM, CNN-GRU, CNN-BiLSTM, and
244 CNN-BiGRU, in terms of inhalation, exhalation, and adventitious sound detection. Fig 4 illustrates
245 the detailed model structures. The outputs of the LSTM, GRU, BiLSTM, and BiGRU models were
246 938×1 vectors, and those of the CNN-LSTM, CNN-GRU, CNN-BiLSTM, and CNN-BiGRU
247 models were 469×1 vectors. An element in these vectors was set to 1 if an inhalation, exhalation,
248 CAS, or DAS occurred within a time segment in which the output value passed the thresholding
249 criterion; otherwise, the element was set to 0.

250

251 **Fig. 4. Model architectures and postprocessing for inhalation, exhalation, CAS, and DAS**
252 **segment and event detection.** (a) LSTM and GRU models; (b) BiLSTM and BiGRU models; and (c)
253 CNN-LSTM, CNN-GRU, CNN-BiLSTM, and CNN-BiGRU models.

254

255 For a fairer comparison of the performance of the unidirectional and bidirectional models, we
256 trained additional simplified (SIMP) BiLSTM, SIMP BiGRU, SIMP CNN-BiLSTM, and SIMP
257 CNN-BiGRU models by adjusting the number of trainable parameters. Parameter adjustment was
258 conducted by halving the number of cells of the LSTM and GRU layers.

259 We used Adam as the optimizer in the benchmark model, and we set the initial learning rate to
260 0.0001 with a step decay ($0.2\times$) when the validation loss did not decrease for 10 epochs. The learning

261 process stopped when no improvement occurred over 50 consecutive epochs.

262

263 *3.4 Postprocessing*

264 The prediction vectors obtained using the adopted models can be further processed for different
265 purposes. For example, we can transform the prediction result from frames to time for real-time
266 monitoring. The breathing duration of most humans lies within a certain range; we considered this
267 fact in our study. Accordingly, when the prediction results obtained using the models indicated that
268 two consecutive inhalation events occurred within a very small interval, we checked the continuity of
269 these two events and decided whether to merge them, as illustrated in the bottom panel of Fig 4a. For
270 example, when the interval between the j th and i th events was smaller than T s, we computed the
271 difference in frequency between their energy peaks ($|\mathbf{p}_j - \mathbf{p}_i|$). Subsequently, if the difference was
272 below a given threshold P , the two events were merged into a single event. In the experiment, T was
273 set to 0.5 s, and P was set to 25 Hz. After the merging process, we further assessed whether a burst
274 event existed. If the duration of an event was shorter than 0.05 s, the event was deleted.

275

276 *3.5 Dataset arrangement and cross-validation*

277 We adopted fivefold cross-validation in the training dataset to train and validate the models.

278 Moreover, we used an independent testing dataset to test the performance of the trained models.

279 According to our preliminary experience, the acoustic patterns of the breath sounds collected from

280 one patient at different auscultation locations or between short intervals had many similarities. To
 281 avoid potential data leakage caused by our methods of collecting and truncating the breath sound
 282 signals, we assigned all truncated recordings collected on the same day to only one of the training,
 283 validation, or testing datasets; this is because these recordings might have been collected from the
 284 same patient within a short period. The statistics of the datasets are listed in Table 3. We used only
 285 audio files containing CASs and DASs to train and test their corresponding detection models.

286

287 **Table 3. Statistics of the datasets and labels of the HF_Lung_V1 database.**

	Training Dataset	Testing Dataset	Total
Recordings			
No of 15-sec recordings	7809	1956	9765
Total duration (min)	1952.25	489	2441.25
Labels			
No of I	27223	6872	34095
Total duration of I (min)	422.17	105.97	528.14
Mean duration of I (sec)	0.93	0.93	0.93
No of E	15601	2748	18349
Total duration of E (min)	248.05	44.81	292.85
Mean duration of E (sec)	0.95	0.98	0.96
No of C/W/S/R	11464/7027/657/3780	2419/1430/29/960	13883/8457/686/4740
Total duration of C/W/S/R (min)	160.16/100.71/9.10/50.35	31.01/19.02/0.36/11.63	191.16/119.73/9.46/61.98
Mean duration of C/W/S/R (sec)	0.84/0.86/0.83/0.80	0.77/0.80/0.74/0.73	0.83/0.85/0.83/0.78
No of D	13794	1812	15606
Total duration of D (min)	203.59	27.29	230.87
Mean duration of D (sec)	0.89	0.90	0.89

288 I: inhalation, E: exhalation, W: wheeze, S: stridor, R: rhonchus, C: continuous adventitious sound, D: discontinuous adventitious sound. W, S, and R were
289 combined to form C.

290

291

292 3.6 Task definition and evaluation metrics

293 [4] clearly defined classification and detection at the segment, event, and recording levels. In
294 this study, we performed two tasks. The first task involved performing detection at the segment level.
295 The acoustic signal of each lung sound recording was transformed into a spectrogram. The temporal
296 resolution of the spectrogram depended on the window size and overlap ratio of the STFT. The
297 aforementioned parameters were fixed such that each spectrogram was a matrix of size 938×129 .
298 Thus, each recording contained 938 time segments (time frames), and each time segment was
299 automatically labeled (Fig 5b) according to the ground-truth event labels (Fig 5a) assigned by the
300 labelers. The output of the prediction process was a sequential prediction matrix (Fig 5c) of size 938
301 $\times 1$ in the LSTM, GRU, BiLSTM, and BiGRU models and size 469×1 in the CNN-LSTM,
302 CNN-GRU, CNN-BiLSTM, and CNN-BiGRU models. By comparing the sequential prediction with
303 the ground-truth time segments, we could define true positive (TP; orange vertical bars in Fig 5d),
304 true negative (TN; green vertical bars in Fig 5d), false positive (FP; black vertical bars in Fig 5d),
305 and false negative (FN; yellow vertical bars in Fig 5d) time segments. Subsequently, the models'
306 sensitivity and specificity in classifying the segments in each recording were computed.

307

308 **Fig 5. Task definition and evaluation metrics.** (a) Ground-truth event labels, (b) ground-truth time
309 segments, (c) AI inference results, (d) segment classification, (e) event detection, and (f) legend. JI:
310 Jaccard index.

311

312 The second task entailed event detection at the recording level. After completing the sequential
313 prediction (Fig 5c), we assembled the time segments associated with the same label into a
314 corresponding event (Fig 5e). We also derived the start and end times of each assembled event. The
315 Jaccard index (JI; [27]) was used to determine whether an AI inference result correctly matched the
316 ground-truth event. For an assembled event to be designated as a TP event (orange horizontal bars in
317 Fig 5e), the corresponding JI value must be greater than 0.5. If the JI was between 0 and 0.5, the
318 assembled event was designated as an FN event (yellow horizontal bars in Fig 5e), and if it was 0,
319 the assembled event was designated as an FP event (black horizontal bars in Fig 5e). A TN event
320 cannot be defined in the task of event detection.

321 The performance of the models was evaluated using the *FI* score, and that of segment detection
322 was evaluated using the receiver operating characteristic (ROC) curve and area under the ROC curve
323 (AUC). In addition, the mean absolute percentage error (MAPE) of event detection was derived. The
324 accuracy, positive predictive value (PPV), sensitivity, specificity, and *FI* score of the models are
325 presented in the section of Supporting information.

326

327 *3.7 Hardware and software*

328 We trained the baseline models on an Ubuntu 18.04 server that was provided by the National
329 Center for High-Performance Computing in Taiwan [Taiwan Computing Cloud (TWCC)] and was
330 equipped with an Intel(R) Xeon(R) Gold 6154 @3.00 GHz CPU with 90 GB RAM. To manage the
331 intensive computation involved in RNN training, we implemented the training module by using the
332 TensorFlow 2.10, CUDA 10, and CuDNN 7 programs to run the NVIDIA Titan V100 card on the
333 TWCC server for GPU acceleration.

334 4 Results

335 4.1 LSTM versus GRU models

336 Table 4 presents the *FI* scores used to compare the eight LSTM- and GRU-based models. When
 337 a CNN was not added, the GRU models outperformed the LSTM models by 0.7%–9.5% in terms of
 338 the *FI* scores. However, the CNN-GRU and CNN-BiGRU models did not outperform the
 339 CNN-LSTM and CNN-BiLSTM models in terms of the *FI* scores (and vice versa).

340

341 **Table 4. Comparison of *FI* scores between LSTM-based models and GRU-based models.**

Models	n of trainable parameters	Inhalation		Exhalation		CASs		DASs	
		<i>FI</i> score		<i>FI</i> score		<i>FI</i> score		<i>FI</i> score	
		Segment Detection	Event Detection	Segment Detection	Event Detection	Segment Detection	Event Detection	Segment Detection	Event Detection
LSTM	300,609	73.9%	76.1%	51.8%	57.0%	15.1%	12.2%	62.6%	59.1%
GRU	227,265	76.2%	78.9%	59.8%	65.6%	24.6%	20.1%	65.9%	62.5%
BiLSTM	732,225	78.1%	84.0%	57.3%	63.9%	19.8%	19.1%	69.6%	70.0%
BiGRU	552,769	80.3%	86.2%	64.1%	70.9%	26.9%	25.6%	70.3%	71.4%
CNN-LSTM	3,448,513	77.6%	81.1%	57.7%	62.1%	45.3%	42.5%	68.8%	64.4%
CNN-GRU	2,605,249	78.4%	82.0%	57.2%	62.0%	51.5%	49.8%	68.0%	64.6%
CNN-BiLSTM	6,959,809	80.6%	86.3%	60.4%	65.6%	47.9%	46.4%	71.2%	70.8%
CNN-BiGRU	5,240,513	80.6%	86.2%	62.2%	68.5%	53.3%	51.6%	70.6%	70.0%

342 The bold values indicate the higher *FI* score between the compared pairs of models.

343

344 According to the ROC curves presented in Fig 6a–d, the GRU-based models outperformed the
 345 LSTM-based models in all compared pairs, except for one pair, in terms of DAS segment detection
 346 (AUC of 0.891 for CNN-BiLSTM vs 0.889 for CNN-BiGRU).

347

348 **Fig. 6. ROC curves for (a) inhalation, (b) exhalation, (c) CAS, and (d) DAS segment detection.**

349 The corresponding AUC values are presented.

350 *4.2 Unidirectional versus bidirectional models*

351 As presented in Table 5, the bidirectional models outperformed their unidirectional counterparts
 352 in all the defined tasks by 0.4%–9.8% in terms of the *FI* scores, even when the bidirectional models
 353 had fewer trainable parameters after model adjustment.

354

355 **Table 5. Comparison of *FI* scores between the unidirectional and bidirectional models.**

Models	n of trainable parameters	Inhalation		Exhalation		CASs		DASs	
		<i>FI</i> score		<i>FI</i> score		<i>FI</i> score		<i>FI</i> score	
		Segment Detection	Event Detection	Segment Detection	Event Detection	Segment Detection	Event Detection	Segment Detection	Event Detection
LSTM	300,609	73.9%	76.1%	51.8%	57.0%	15.1%	12.2%	62.6%	59.1%
SIMP BiLSTM	235,073	77.8%	84.1%	55.8%	62.4%	19.8%	17.9%	68.8%	68.9%
GRU	227,265	76.2%	78.9%	59.8%	65.6%	24.6%	20.1%	65.9%	62.5%
SIMP BiGRU	178,113	80.1%	86.1%	63.7%	70.0%	25.0%	22.2%	70.3%	71.3%
CNN-LSTM	3,448,513	77.6%	81.1%	57.7%	62.1%	45.3%	42.5%	68.8%	64.4%
SIMP CNN-BiLSTM	3,382,977	80.0%	85.8%	60.4%	66.2%	50.8%	50.2%	70.2%	70.2%
CNN-GRU	2,605,249	78.4%	82.0%	57.2%	62.0%	51.5%	49.8%	68.0%	64.6%
SIMP CNN-BiGRU	2,556,097	80.1%	85.9%	62.4%	68.4%	52.6%	51.5%	69.9%	69.5%

356 The bold values indicate the higher *FI* score between the compared pairs of models. SIMP means the number of trainable parameters
 357 is adjusted.

358

359 *4.3 Models with CNN versus those without CNN*

360 According to Table 6, the models with a CNN outperformed those without a CNN in 26 of the

361 32 compared pairs.

362

363

364 **Table 6. Comparison of *FI* scores between models without and with a CNN.**

Models	n of trainable parameters	Inhalation		Exhalation		CASs		DASs	
		<i>FI</i> score		<i>FI</i> score		<i>FI</i> score		<i>FI</i> score	
		Segment Detection	Event Detection	Segment Detection	Event Detection	Segment Detection	Event Detection	Segment Detection	Event Detection
LSTM	300,609	73.9%	76.1%	51.8%	57.0%	15.10%	12.20%	62.60%	59.10%
CNN-LSTM	3,448,513	77.6%	81.1%	57.7%	62.1%	45.30%	42.50%	68.80%	64.40%
BiLSTM	732,225	76.2%	78.9%	59.8%	65.6%	19.80%	17.90%	68.80%	68.90%
CNN-BiLSTM	6,959,809	78.4%	82.0%	57.2%	62.0%	50.80%	50.20%	70.20%	70.20%
GRU	227,265	78.1%	84.0%	57.3%	63.9%	24.60%	20.10%	65.90%	62.50%
CNN-GRU	2,605,249	80.6%	86.3%	60.4%	65.6%	51.50%	49.80%	68.00%	64.60%
BiGRU	178,113	80.3%	86.2%	64.1%	70.9%	25.00%	22.20%	70.30%	71.30%
CNN-BiGRU	2,556,097	80.6%	86.2%	62.2%	68.5%	52.60%	51.50%	69.90%	69.50%

365 The bold values indicate the higher *FI* score between the compared pairs of models.

366

367 The models with a CNN exhibited higher AUC values than did those without a CNN (Fig 6a–d),
368 except that BiGRU had a higher AUC value than did CNN-BiGRU in terms of inhalation detection
369 (0.963 vs 0.961), GRU had a higher AUC value than did CNN-GRU in terms of exhalation detection
370 (0.886 vs 0.883), and BiGRU had a higher AUC value than did CNN-BiGRU in terms of exhalation
371 detection (0.911 vs 0.899).

372 Moreover, compared with the LSTM, GRU, BiLSTM, and BiGRU models, the CNN-LSTM,

373 CNN-GRU, CNN-BiLSTM, and CNN-BiGRU models exhibited flatter and lower MAPE curves

374 over a wide range of threshold values in all event detection tasks (Fig 7a–d).

375

376

377 **Fig 7. MAPE curves for (a) inhalation, (b) exhalation, (c) CAS, and (d) DAS event detection.**

378 **5 Discussion**

379 *5.1 Benchmark results*

380 According to the *F1* scores presented in Table 4, among models without a CNN, the GRU and
381 BiGRU models consistently outperformed the LSTM and BiLSTM models in all defined tasks.
382 However, the GRU-based models did not have superior *F1* scores among models with a CNN.
383 Regarding the ROC curves and AUC values (Fig 6a–d), the GRU-based models consistently
384 outperformed the other models in all but one task. Accordingly, we can conclude that GRU-based
385 models perform slightly better than LSTM-based models in lung sound analysis. Previous studies
386 have also compared LSTM- and GRU-based models [38, 46, 47]. Although a concrete conclusion
387 cannot be drawn regarding whether LSTM-based models are superior to the GRU-based models (and
388 vice versa), GRU-based models have been reported to outperform LSTM-based models in terms of
389 computation time [38, 47].

390 As presented in Table 5, the bidirectional models outperformed their unidirectional counterparts
391 in all defined tasks, a finding that is consistent with several previously obtained results [29, 36, 38,
392 40].

393 A CNN can facilitate the extraction of useful features and enhance the prediction accuracy of
394 RNN-based models. The benefits engendered by a CNN are particularly vital in CAS detection. For
395 the models with a CNN, the *FI* score improvement ranged from 26.0% to 30.3% and the AUC
396 improvement ranged from 0.067 to 0.089 in the CAS detection tasks. Accordingly, we can infer that
397 considerable information used in CAS detection resides in the local positional arrangement of the
398 features. Thus, a two-dimensional CNN facilitates the extraction of the associated information.
399 Notably, CNN-induced improvements in model performance in the inhalation, exhalation, and DAS
400 detection tasks were not as high as those observed in the CAS detection tasks. The MAPE curves
401 (Fig 7a–d) reveal that a model with a CNN has more consistent predictions over various threshold
402 values.

403 In our previous study [26], an attention-based encoder–decoder architecture based on ResNet
404 and LSTM exhibited favorable performance in inhalation (*FI* score of 90.4%) and exhalation (*FI*
405 score of 93.2%) segment detection tasks. However, the model was established on the basis of a very
406 small dataset (489 recordings of 15-s-long lung sounds). Moreover, the model involves a
407 complicated architecture; hence, it is impossible to implement real-time respiratory monitoring in
408 devices with limited computing power, such as smartphones or medical-grade tablets.

409 Few studies have performed event detection at the recording level by using a comparatively
410 simple deep learning model. [29] used the BiGRU model and one-dimensional labels (similar to
411 those used in the present study) for breath phase and crackle detection. Their BiGRU model

412 exhibited comparable performance to our models in terms of inhalation event detection (*FI* scores,
413 87.0% vs 86.2%) and in terms of DAS event detection (*FI* scores, 72.1% vs 71.4%). However, the
414 performance of the BiGRU model differed considerably from that of our models in terms of
415 exhalation detection (*FI* scores: 84.6% vs 70.9%). One of the reasons for this discrepancy is that [29]
416 established their ground-truth labels on the basis of the gold-standard signals of a pneumotachograph.
417 Another reason is that an exhalation label is not always available following an inhalation label in our
418 data. Finally, we did not specifically control the sounds we recorded; for example, we did not ask
419 patients to perform voluntary deep breathing or keep ambient noise down. The factors influencing
420 the model performance are further discussed in the next section.

421

422 5.2 *Factors influencing model performance*

423 The benchmark performance of the proposed models may have been influenced by the
424 following factors: (1) unusual breathing patterns; (2) imbalanced data; (3) low signal-to-noise ratio
425 (SNR); (4) noisy labels, including class and attribute noise, in the database; and (5) sound
426 overlapping.

427 Fig 8 displays most of the breath patterns present in the HF_Lung_V1 database. Fig 8a
428 illustrates the general pattern of a breath cycle in the lung sounds when the ratio of inhalation to
429 exhalation durations is approximately 2:1 and an expiratory pause is noted [3, 4]. Fig 8b presents a
430 frequent condition under which an exhalation is not completely heard by the labelers. However,

431 because we did not ask the subjects to breath voluntarily when recording the sound, many unusual
432 breath patterns might have been recorded, such as patterns caused by shallow breathing, fast
433 breathing, and apnea as well as those caused by double triggering of the ventilator [48] and air
434 trapping [49, 50]. These unusual breathing patterns might confuse the labeling and learning
435 processes and result in poor testing results.

436

437 **Fig 8. Patterns of normal breathing lung sounds.** (a) General lung sound patterns and (b) general
438 lung sound patterns with unidentifiable exhalations. “I” represents an identifiable inhalation event, “E”
439 represents an identifiable exhalation event, and the black areas represent pause phases.

440

441 The developed database contains imbalanced numbers of inhalation and exhalation labels
442 (34,095 and 18,349, respectively) because not every exhalation was heard and labeled. In addition,
443 the proposed models may possess the capability of learning the rhythmic rise and fall of breathing
444 signals but not the capability of learning acoustic or texture features that can distinguish an
445 inhalation from an exhalation. This may thus explain the models’ poor performance in exhalation
446 detection. However, these models are suitable for respiratory rate estimation and apnea detection as
447 long as appropriate inhalation detection is achieved. Furthermore, for all labels, the summation of the
448 event duration was smaller than that of the background signal duration (these factors had a ratio of
449 approximately 1:2.5 to 1:7). The aforementioned phenomenon can be regarded as foreground–
450 background class imbalance [51] and will be addressed in future studies.

451 Most of the sounds in the established database were not recorded during the patients performed
452 deep breathing; thus, the signal quality was not maximized. However, training models with such
453 nonoptimal data increase their adaptability to real-world scenarios. Moreover, the SNR may be
454 reduced by noise, such as human voices; music; sounds from bedside monitors, televisions, air
455 conditioners, fans, and radios; sounds generated by mechanical ventilators; electrical noise generated
456 by touching or moving the parts of acoustic sensors; and friction sounds generated by the rubbing of
457 two surfaces together (e.g., rubbing clothes with the skin). A poor SNR of audio signals can lead to
458 difficulties in labeling and prediction tasks. The features of some noise types are considerably similar
459 to those of adventitious sounds. The poor performance of the proposed models in CAS detection can
460 be partly attributed to the noisy environment in which the lung sounds were recorded. In particular,
461 the sounds generated by ventilators caused numerous FP events in the CAS detection tasks. Thus,
462 additional effort is required to develop a superior preprocessing algorithm that can filter out
463 influential noise or to identify a strategy to ensure that models focus on learning the correct CAS
464 features. Furthermore, the integration of active noise-canceling technology [52] or noise suppression
465 technology [53] into respiratory sound monitors can help reduce the noise from auscultatory signals.

466 The sound recordings in the HF_Lung_V1 database were labeled by only one labeler; thus,
467 some noisy labels, including class and attribute noise, may exist in the database [54]. These noisy
468 labels are attributable to (1) the different hearing abilities of the labeler, which can cause differences
469 in the labeled duration; (2) the absence of clear criteria for differentiating between target and

470 confusing events; (3) individual human errors; (4) tendency to not label events located close to the
471 beginning and end of a recording; and (5) confusion caused by unusual breath patterns and poor
472 SNRs. However, deep learning models exhibit high robustness to noisy labels [55]. Accordingly, we
473 are currently working toward establishing better ground-truth labels.

474 Breathing generates CASs and DASs under abnormal respiratory conditions. This means that
475 the breathing sound, CAS, and DAS might overlap with one another during the same period. This
476 sound overlapping, along with the data imbalance, makes the CAS and DAS detection models learn
477 to read the rise and fall of the breathing energy and falsely identify an inhalation or exhalation as
478 CAS or DAS, respectively. This FP detection was observed in our benchmark results. In the future,
479 strategies must be adopted to address the problem of sound overlap.

480

481 **6 Conclusions**

482 We established a large open-access lung sound database, namely HF_Lung_V1
483 (https://gitlab.com/techsupportHF/HF_Lung_V1), that contains 9,765 audio files of lung sounds
484 (each with a duration of 15 s), 34,095 inhalation labels, 18,349 exhalation labels, 13,883 CAS labels
485 (comprising 8,457 wheeze labels, 686 stridor labels, and 4,740 rhonchus labels), and 15,606 DAS
486 labels (all of which are crackles).

487 We also investigated the performance of eight RNN-based models in terms of inhalation,
488 exhalation, CAS detection, and DAS detection in the HF_Lung_V1 database. We determined that the

489 bidirectional models outperformed the unidirectional models in lung sound analysis. Furthermore,
490 the addition of a CNN to these models further improved their performance.

491 Future studies can develop more accurate respiratory sound analysis models. First, highly
492 accurate ground-truth labels should be established. Second, researchers should investigate the
493 performance of RNN-based models containing state-of-the-art convolutional layers. Third, regional
494 CNN variants can be adopted in lung sound analysis if the labels are expanded to two-dimensional
495 bounding boxes [27]. Fourth, wavelet-based approaches, empirical mode decomposition, and other
496 methods that can extract different features should be investigated [4, 56]. Finally, respiratory sound
497 monitors should be equipped with the capability of tracheal breath sound analysis [52].

498

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505

- 507 1. Bohadana A, Izbicki G, Kraman SS. Fundamentals of lung auscultation. *New England Journal*
508 *of Medicine*. 2014;370(8):744-51.
- 509 2. Goettel N, Herrmann MJ. *Breath Sounds: From Basic Science to Clinical Practice*. *Anesthesia*
510 *& Analgesia*. 2019;128(3):e42.
- 511 3. Sarkar M, Madabhavi I, Niranjana N, Dogra M. Auscultation of the respiratory system. *Annals*
512 *of thoracic medicine*. 2015;10(3):158.
- 513 4. Pramono RXA, Bowyer S, Rodriguez-Villegas E. Automatic adventitious respiratory sound
514 analysis: A systematic review. *PloS one*. 2017;12(5):e0177926.
- 515 5. Wang B, Liu Y, Wang Y, Yin W, Liu T, Liu D, et al. Characteristics of Pulmonary auscultation
516 in patients with 2019 novel coronavirus in china. 2020.
- 517 6. Raj V, Renjini A, Swapna M, Sreejyothi S, Sankararaman S. Nonlinear time series and principal
518 component analyses: Potential diagnostic tools for COVID-19 auscultation. *Chaos, Solitons &*
519 *Fractals*. 2020;140:110246.
- 520 7. Sovijärvi A, Vanderschoot J, Earis J. Standardization of computerized respiratory sound
521 analysis. *Crit Care Med*. 1997;156:974-87.
- 522 8. Berry MP, Martí J-D, Ntoumenopoulos G. Inter-rater agreement of auscultation, palpable
523 fremitus, and ventilator waveform sawtooth patterns between clinicians. *Respiratory care*.
524 2016;61(10):1374-83.
- 525 9. Grunnreis FO. Intra-and interobserver variation in lung sound classification. Effect of training:
526 UiT Norges arktiske universitet; 2016.
- 527 10. Gurung A, Scrafford CG, Tielsch JM, Levine OS, Checkley W. Computerized lung sound
528 analysis as diagnostic aid for the detection of abnormal lung sounds: a systematic review and
529 meta-analysis. *Respiratory medicine*. 2011;105(9):1396-403.
- 530 11. Huq S, Moussavi Z. Acoustic breath-phase detection using tracheal breath sounds. *Medical &*
531 *biological engineering & computing*. 2012;50(3):297-308.
- 532 12. Mesaros A, Heittola T, Virtanen T. Metrics for polyphonic sound event detection. *Applied*
533 *Sciences*. 2016;6(6):162.
- 534 13. Pasterkamp H, Kraman SS, Wodicka GR. Respiratory sounds: advances beyond the
535 stethoscope. *American journal of respiratory and critical care medicine*. 1997;156(3):974-87.
- 536 14. Chambres G, Hanna P, Desainte-Catherine M, editors. Automatic detection of patient with
537 respiratory diseases using lung sound analysis. 2018 International Conference on Content-Based
538 Multimedia Indexing (CBMI); 2018: IEEE.
- 539 15. Demir F, Sengur A, Bajaj V. Convolutional neural networks based efficient approach for
540 classification of lung diseases. *Health Inf Sci Syst*. 2020;8(1):4.

- 541 16. Hosseini M, Ren H, Rashid H-A, Mazumder AN, Prakash B, Mohsenin T. Neural Networks for
542 Pulmonary Disease Diagnosis using Auditory and Demographic Information. arXiv preprint
543 arXiv:201113194. 2020.
- 544 17. Perna D, Tagarelli A. Deep Auscultation: Predicting Respiratory Anomalies and Diseases via
545 Recurrent Neural Networks. 2019 IEEE 32nd International Symposium on Computer-Based
546 Medical Systems (CBMS)2019. p. 50-5.
- 547 18. Pham L, McLoughlin I, Phan H, Tran M, Nguyen T, Palaniappan R. Robust Deep Learning
548 Framework For Predicting Respiratory Anomalies and Diseases. arXiv preprint arXiv:200203894.
549 2020.
- 550 19. Acharya J, Basu A. Deep Neural Network for Respiratory Sound Classification in Wearable
551 Devices Enabled by Patient Specific Model Tuning. IEEE transactions on biomedical circuits and
552 systems. 2020;14(3):535-44.
- 553 20. Aykanat M, Kılıç Ö, Kurt B, Saryal S. Classification of lung sounds using convolutional neural
554 networks. EURASIP Journal on Image and Video Processing. 2017;2017(1).
- 555 21. Bardou D, Zhang K, Ahmad SM. Lung sounds classification using convolutional neural
556 networks. Artif Intell Med. 2018;88:58-69.
- 557 22. Chen H, Yuan X, Pei Z, Li M, Li J. Triple-Classification of Respiratory Sounds Using
558 Optimized S-Transform and Deep Residual Networks. IEEE Access. 2019;7:32845-52.
- 559 23. Grzywalski T, Piecuch M, Szajek M, Breborowicz A, Hafke-Dys H, Kocinski J, et al. Practical
560 implementation of artificial intelligence algorithms in pulmonary auscultation examination. Eur J
561 Pediatr. 2019;178(6):883-90.
- 562 24. Kochetov K, Putin E, Balashov M, Filchenkov A, Shalyto A. Noise Masking Recurrent Neural
563 Network for Respiratory Sound Classification. Artificial Neural Networks and Machine Learning –
564 ICANN 2018. Lecture Notes in Computer Science2018. p. 208-17.
- 565 25. Li L, Xu W, Hong Q, Tong F, Wu J, editors. Classification between normal and adventitious
566 lung sounds using deep neural network. 2016 10th International Symposium on Chinese Spoken
567 Language Processing (ISCSLP); 2016: IEEE.
- 568 26. Hsiao C-H, Lin T-W, Lin C-W, Hsu F-S, Lin FY-S, Chen C-W, et al., editors. Breathing Sound
569 Segmentation and Detection Using Transfer Learning Techniques on an Attention-Based
570 Encoder-Decoder Architecture. 2020 42nd Annual International Conference of the IEEE Engineering
571 in Medicine & Biology Society (EMBC); 2020: IEEE.
- 572 27. Jácome C, Ravn J, Holsbø E, Aviles-Solis JC, Melbye H, Ailo Bongo L. Convolutional neural
573 network for breathing phase detection in lung sounds. Sensors. 2019;19(8):1798.
- 574 28. Liu Y, Lin Y, Gao S, Zhang H, Wang Z, Gao Y, et al., editors. Respiratory sounds feature
575 learning with deep convolutional neural networks. 2017 IEEE 15th Intl Conf on Dependable,
576 Autonomic and Secure Computing, 15th Intl Conf on Pervasive Intelligence and Computing, 3rd Intl

577 Conf on Big Data Intelligence and Computing and Cyber Science and Technology Congress
578 (DASC/PiCom/DataCom/CyberSciTech); 2017: IEEE.

579 29. Messner E, Fediuk M, Swatek P, Scheidl S, Smolle-Juttner F-M, Olschewski H, et al., editors.
580 Crackle and breathing phase detection in lung sounds with deep bidirectional gated recurrent neural
581 networks. 2018 40th Annual International Conference of the IEEE Engineering in Medicine and
582 Biology Society (EMBC); 2018: IEEE.

583 30. Rocha B, Filos D, Mendes L, Vogiatzis I, Perantoni E, Kaimakamis E, et al., editors. A
584 respiratory sound database for the development of automated classification. International Conference
585 on Biomedical and Health Informatics; 2017: Springer.

586 31. Hestness J, Narang S, Ardalani N, Diamos G, Jun H, Kianinejad H, et al. Deep learning scaling
587 is predictable, empirically. arXiv preprint arXiv:171200409. 2017.

588 32. Sun C, Shrivastava A, Singh S, Gupta A, editors. Revisiting unreasonable effectiveness of data
589 in deep learning era. Proceedings of the IEEE international conference on computer vision; 2017.

590 33. Elman JL. Finding structure in time. *Cognitive science*. 1990;14(2):179-211.

591 34. Hochreiter S, Schmidhuber J. Long short-term memory. *Neural computation*.
592 1997;9(8):1735-80.

593 35. Cho K, Van Merriënboer B, Gulcehre C, Bahdanau D, Bougares F, Schwenk H, et al. Learning
594 phrase representations using RNN encoder-decoder for statistical machine translation. arXiv preprint
595 arXiv:14061078. 2014.

596 36. Graves A, Schmidhuber J. Framewise phoneme classification with bidirectional LSTM and
597 other neural network architectures. *Neural networks*. 2005;18(5-6):602-10.

598 37. Schuster M, Paliwal KK. Bidirectional recurrent neural networks. *IEEE transactions on Signal*
599 *Processing*. 1997;45(11):2673-81.

600 38. Khandelwal S, Lecouteux B, Besacier L. Comparing GRU and LSTM for automatic speech
601 recognition. 2016.

602 39. Li L, Wu Z, Xu M, Meng HM, Cai L, editors. Combining CNN and BLSTM to Extract Textual
603 and Acoustic Features for Recognizing Stances in Mandarin Ideological Debate Competition.
604 INTERSPEECH; 2016.

605 40. Parascandolo G, Huttunen H, Virtanen T, editors. Recurrent neural networks for polyphonic
606 sound event detection in real life recordings. 2016 IEEE International Conference on Acoustics,
607 Speech and Signal Processing (ICASSP); 2016: IEEE.

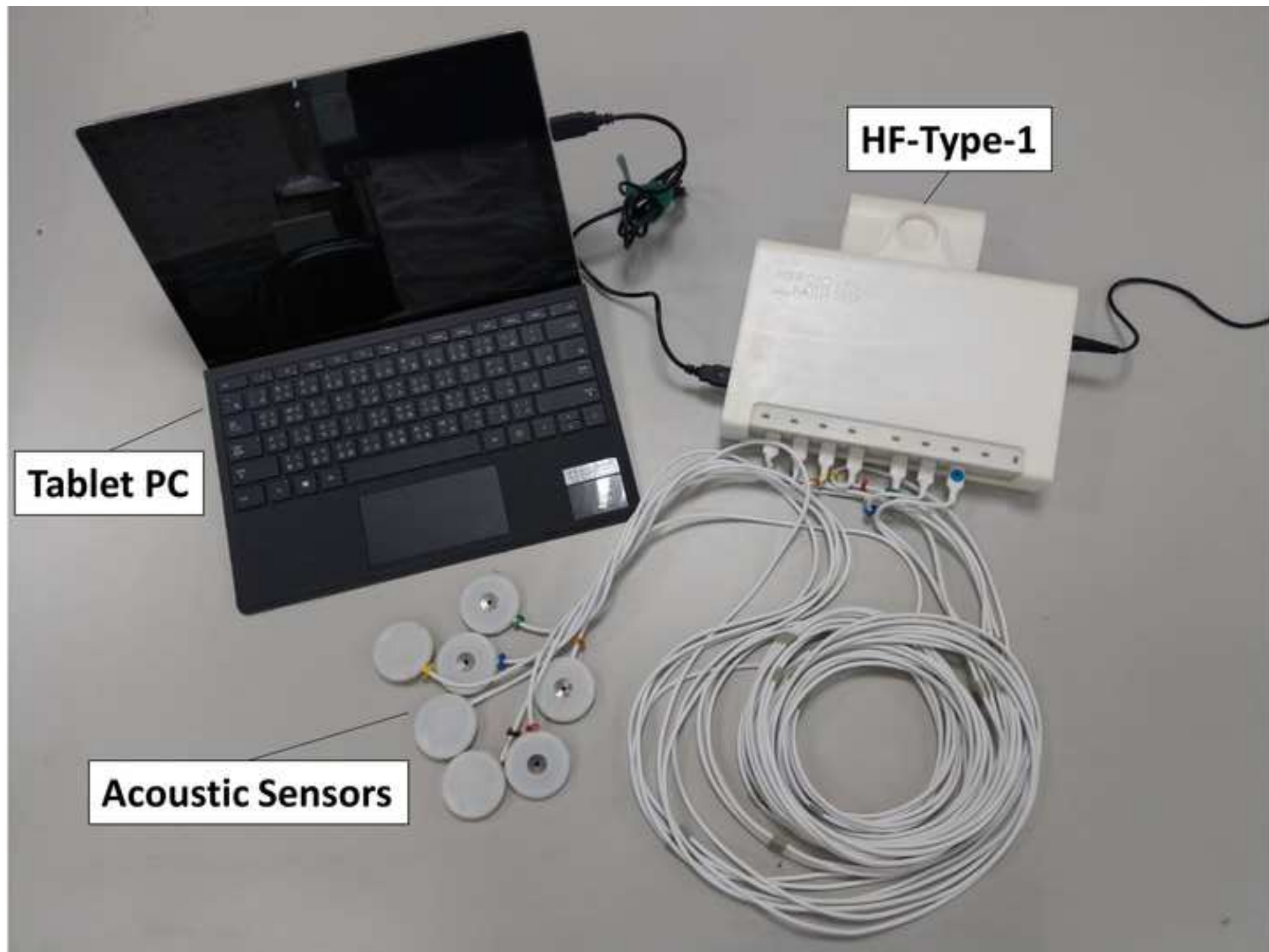
608 41. Deng M, Meng T, Cao J, Wang S, Zhang J, Fan H. Heart sound classification based on
609 improved MFCC features and convolutional recurrent neural networks. *Neural Networks*. 2020.

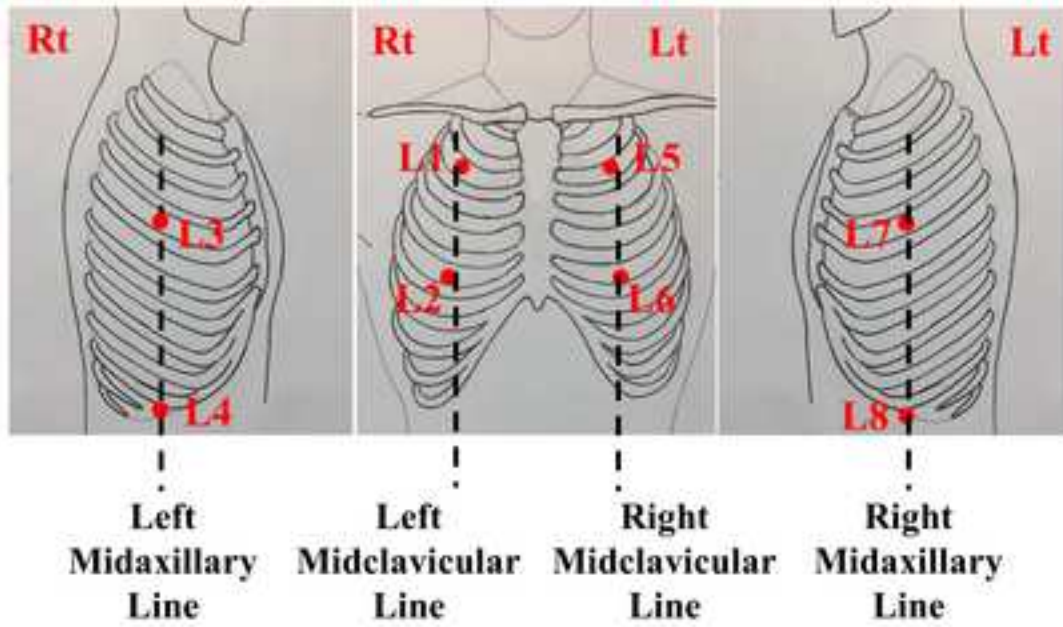
610 42. Zhao H, Zarar S, Tashev I, Lee C-H, editors. Convolutional-recurrent neural networks for
611 speech enhancement. 2018 IEEE International Conference on Acoustics, Speech and Signal
612 Processing (ICASSP); 2018: IEEE.

- 613 43. Pasterkamp H, Brand PL, Everard M, Garcia-Marcos L, Melbye H, Priftis KN. Towards the
614 standardisation of lung sound nomenclature. *European Respiratory Journal*. 2016;47(3):724-32.
- 615 44. Hsu F-S, Huang C-J, Kuo C-Y, Huang S-R, Cheng Y-R, Wang J-H, et al. Development of a
616 respiratory sound labeling software for training a deep learning-based respiratory sound analysis
617 model. arXiv preprint arXiv:210101352. 2021(,).
- 618 45. Chamberlain D, Kodgule R, Ganelin D, Miglani V, Fletcher RR, editors. Application of
619 semi-supervised deep learning to lung sound analysis. 2016 38th Annual International Conference of
620 the IEEE Engineering in Medicine and Biology Society (EMBC); 2016: IEEE.
- 621 46. Chung J, Gulcehre C, Cho K, Bengio Y. Empirical evaluation of gated recurrent neural
622 networks on sequence modeling. arXiv preprint arXiv:14123555. 2014.
- 623 47. Shewalkar AN. Comparison of rnn, lstm and gru on speech recognition data. 2018.
- 624 48. Thille AW, Rodriguez P, Cabello B, Lellouche F, Brochard L. Patient-ventilator asynchrony
625 during assisted mechanical ventilation. *Intensive care medicine*. 2006;32(10):1515-22.
- 626 49. Blanch L, Bernabé F, Lucangelo U. Measurement of air trapping, intrinsic positive
627 end-expiratory pressure, and dynamic hyperinflation in mechanically ventilated patients. *Respiratory
628 care*. 2005;50(1):110-24.
- 629 50. Miller WT, Chatzkel J, Hewitt MG. Expiratory air trapping on thoracic computed tomography.
630 A diagnostic subclassification. *Annals of the American Thoracic Society*. 2014;11(6):874-81.
- 631 51. Oksuz K, Cam BC, Kalkan S, Akbas E. Imbalance problems in object detection: A review.
632 *IEEE Transactions on Pattern Analysis and Machine Intelligence*. 2020.
- 633 52. Wu Y, Liu J, He B, Zhang X, Yu L. Adaptive Filtering Improved Apnea Detection Performance
634 Using Tracheal Sounds in Noisy Environment: A Simulation Study. *BioMed Research International*.
635 2020;2020.
- 636 53. Emmanouilidou D, McCollum ED, Park DE, Elhilali M. Computerized lung sound screening
637 for pediatric auscultation in noisy field environments. *IEEE Transactions on Biomedical
638 Engineering*. 2017;65(7):1564-74.
- 639 54. Zhu X, Wu X. Class noise vs. attribute noise: A quantitative study. *Artificial intelligence
640 review*. 2004;22(3):177-210.
- 641 55. Rolnick D, Veit A, Belongie S, Shavit N. Deep learning is robust to massive label noise. arXiv
642 preprint arXiv:170510694. 2017.
- 643 56. Pramono RXA, Imtiaz SA, Rodriguez-Villegas E. Evaluation of features for classification of
644 wheezes and normal respiratory sounds. *PloS one*. 2019;14(3):e0213659.

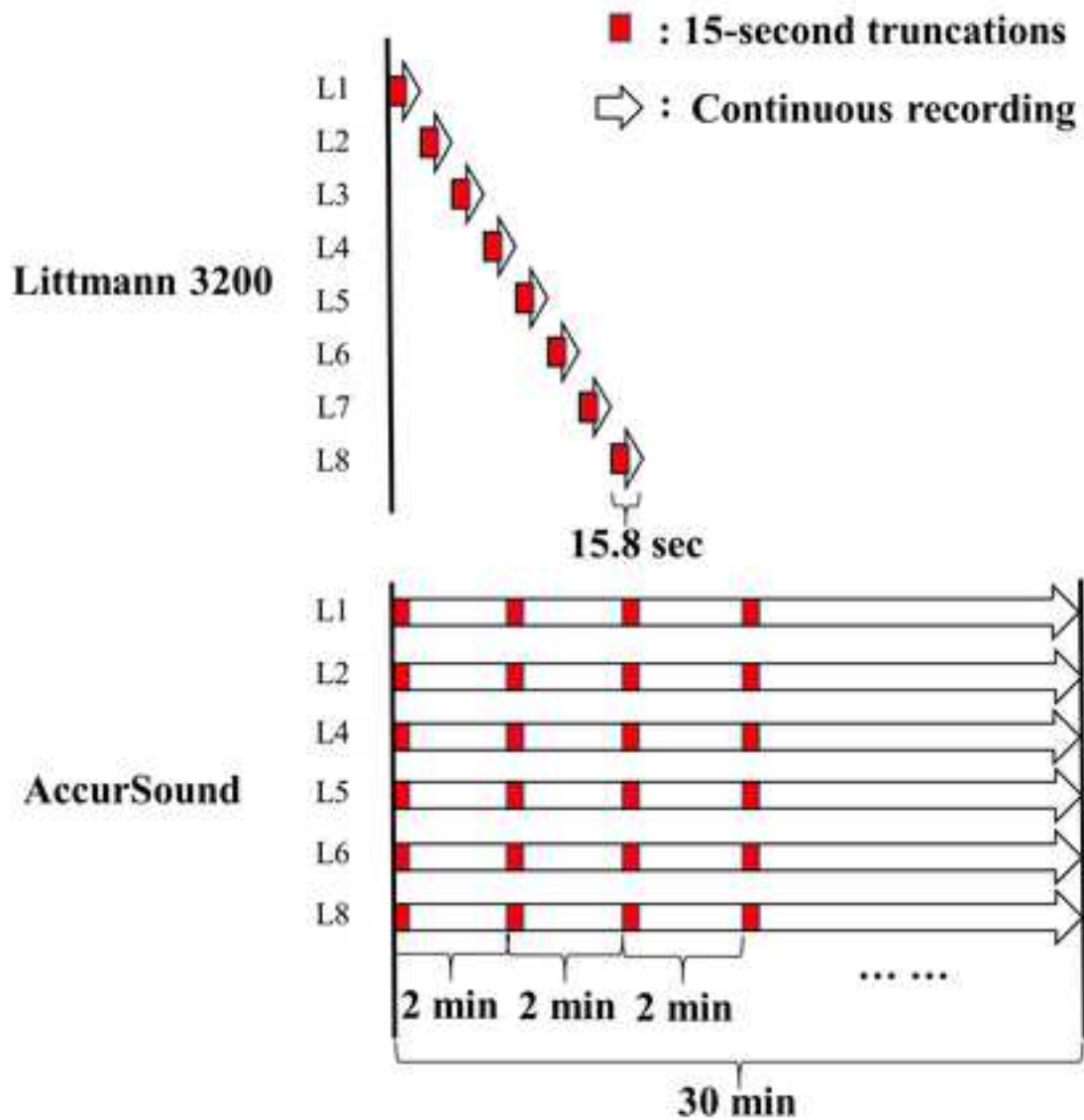
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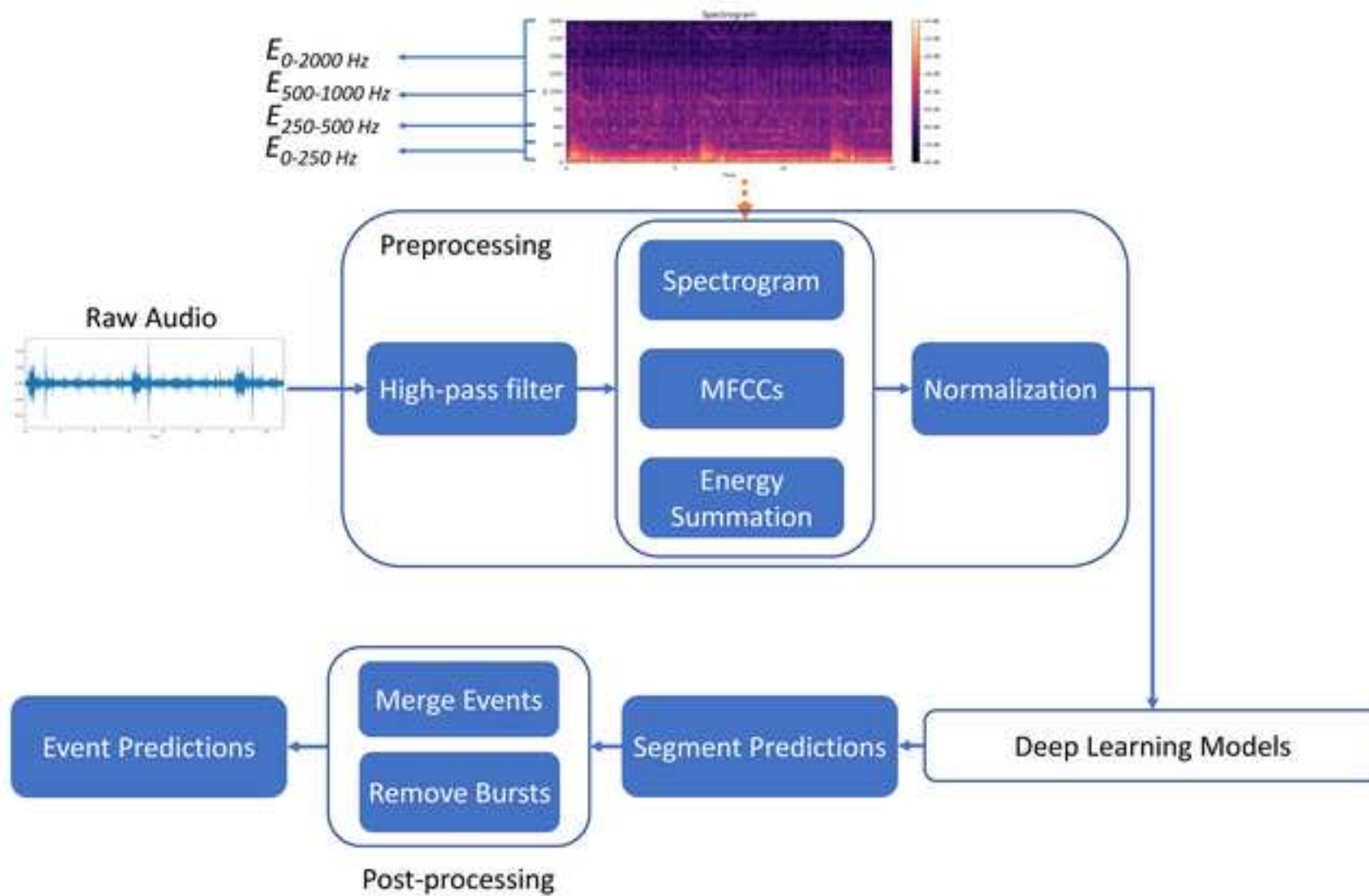


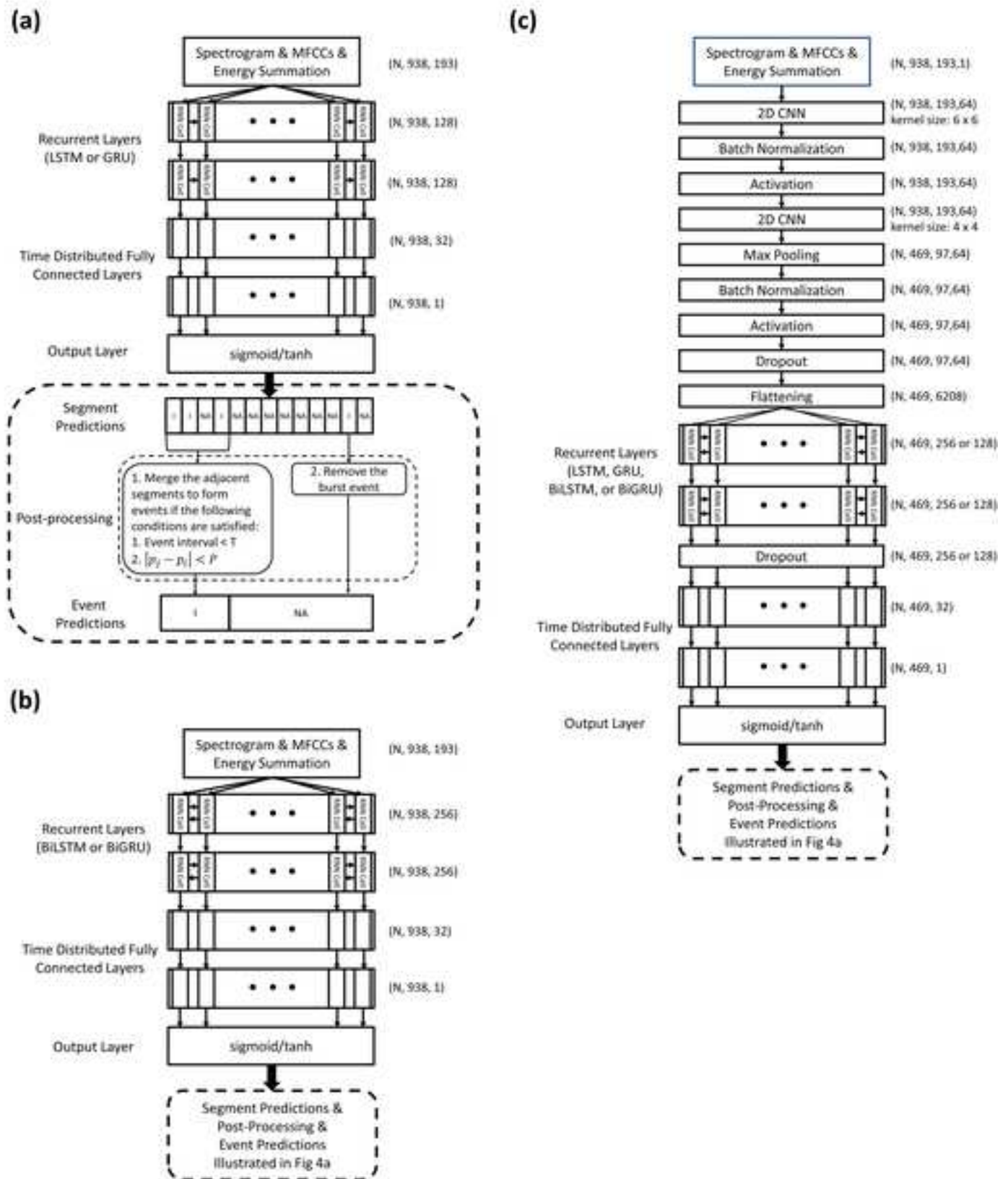


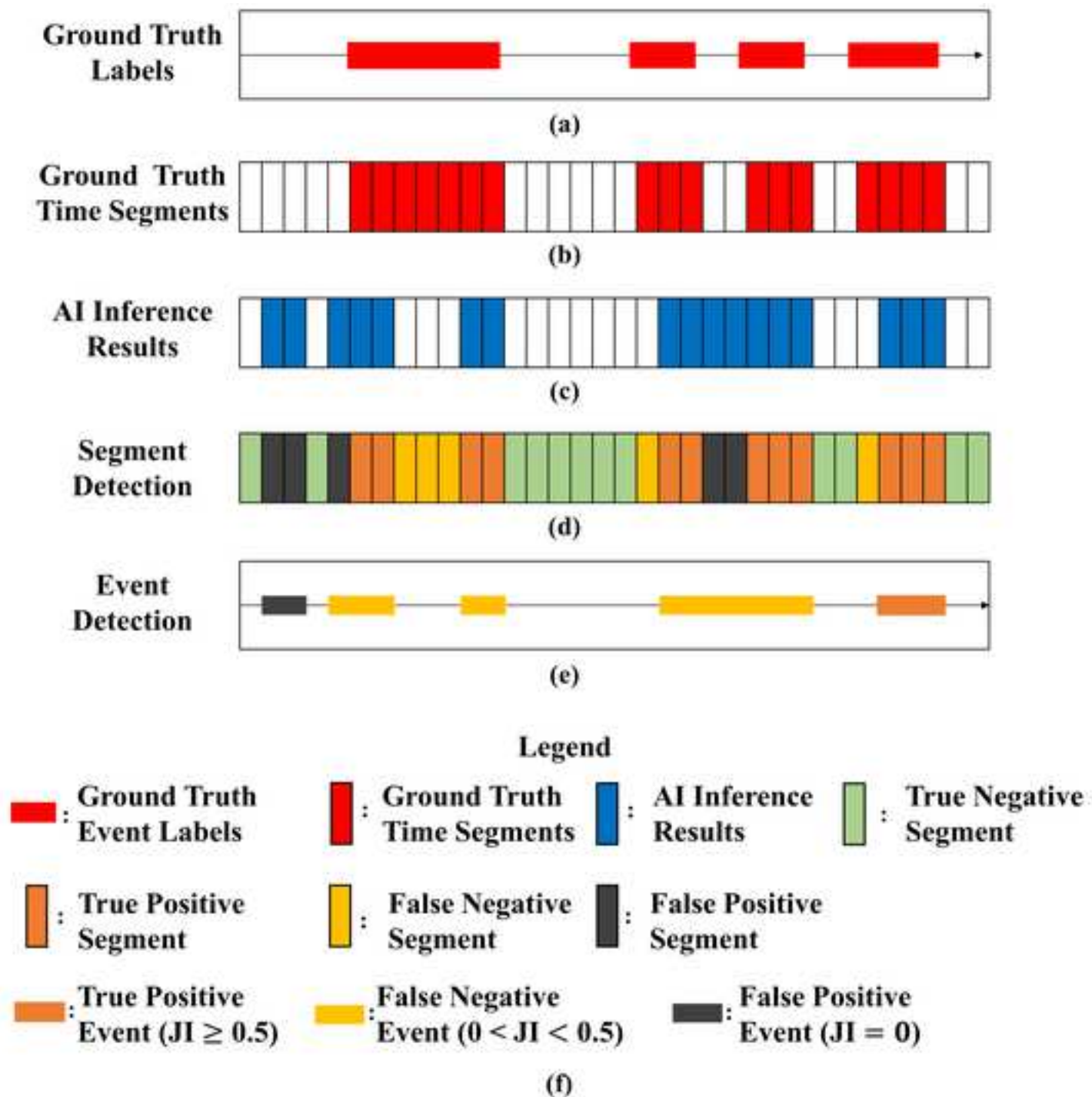
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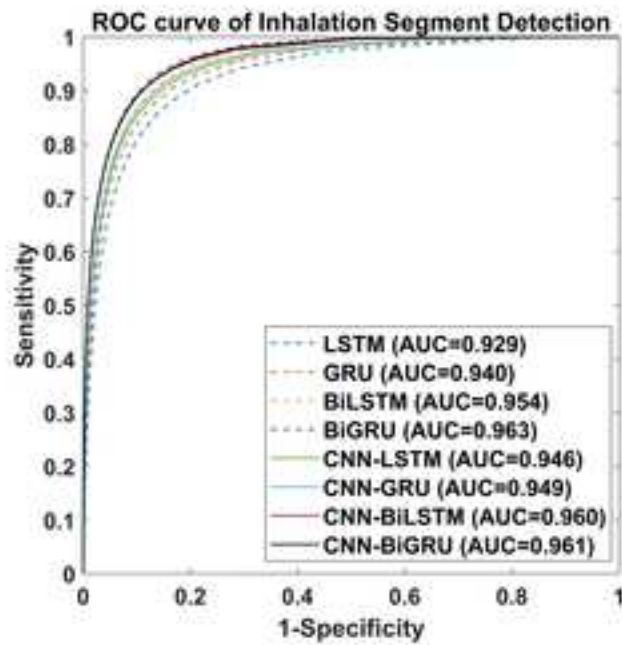


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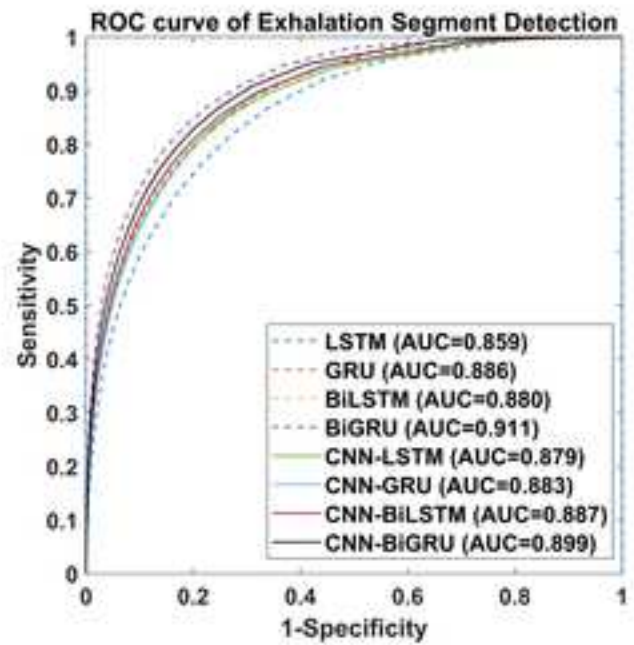




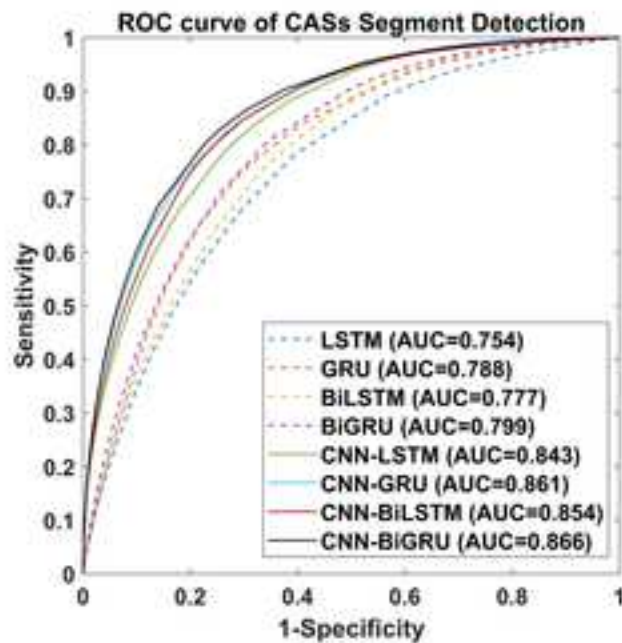




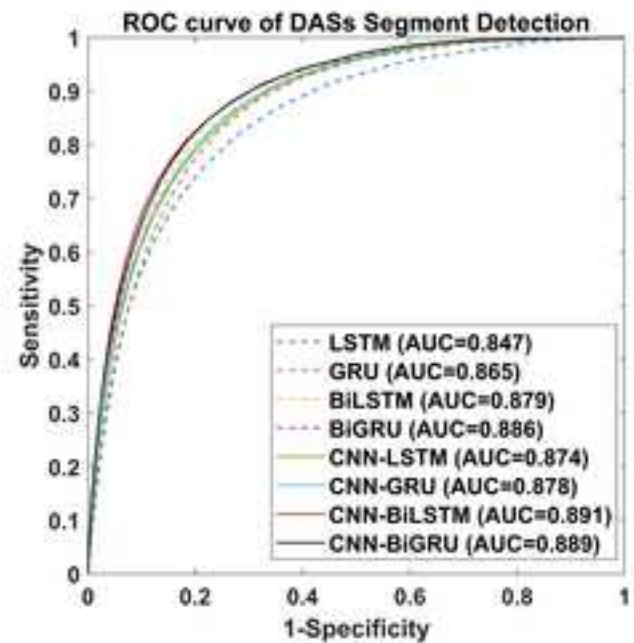
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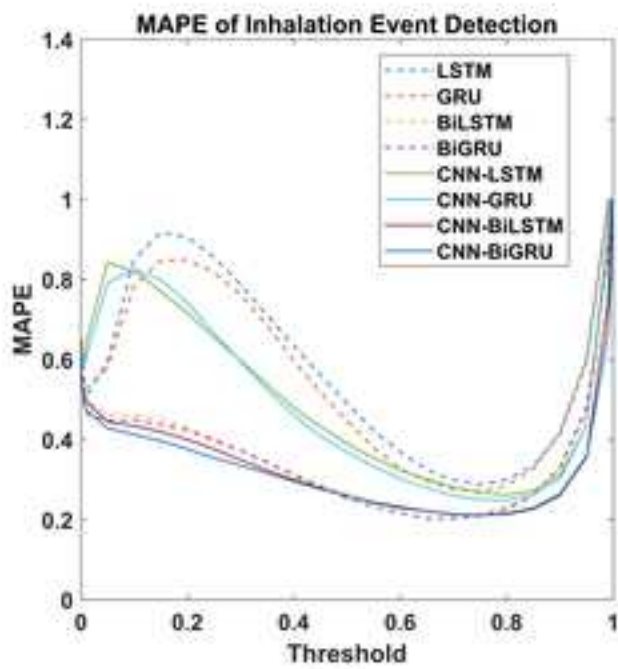
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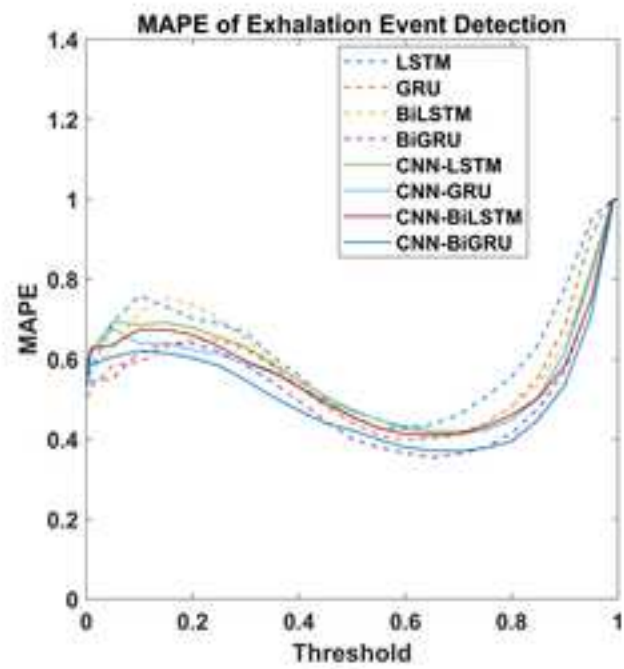
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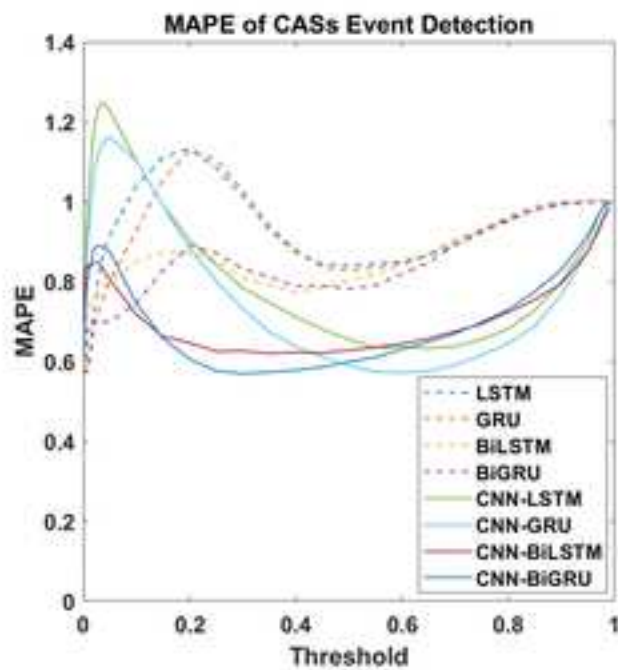
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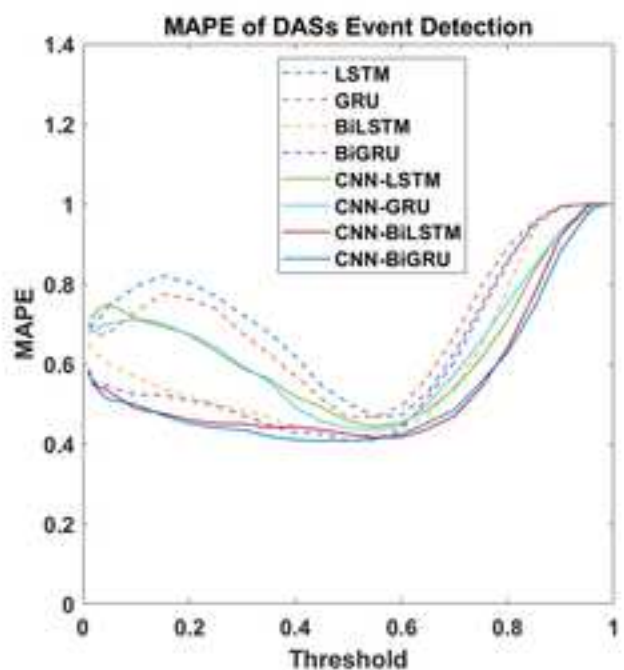
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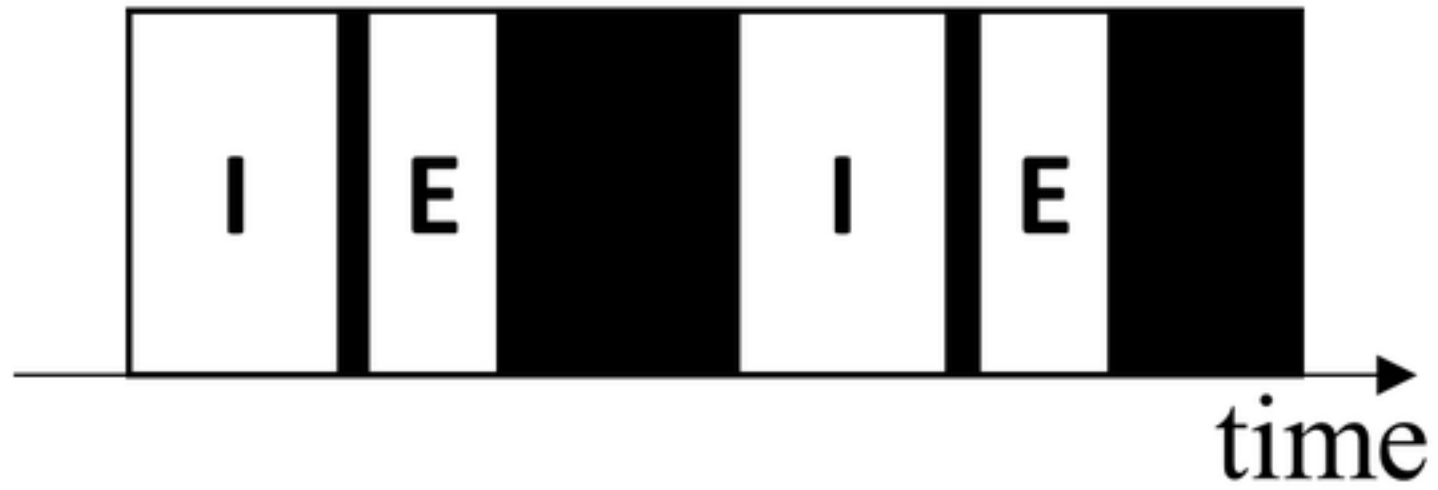
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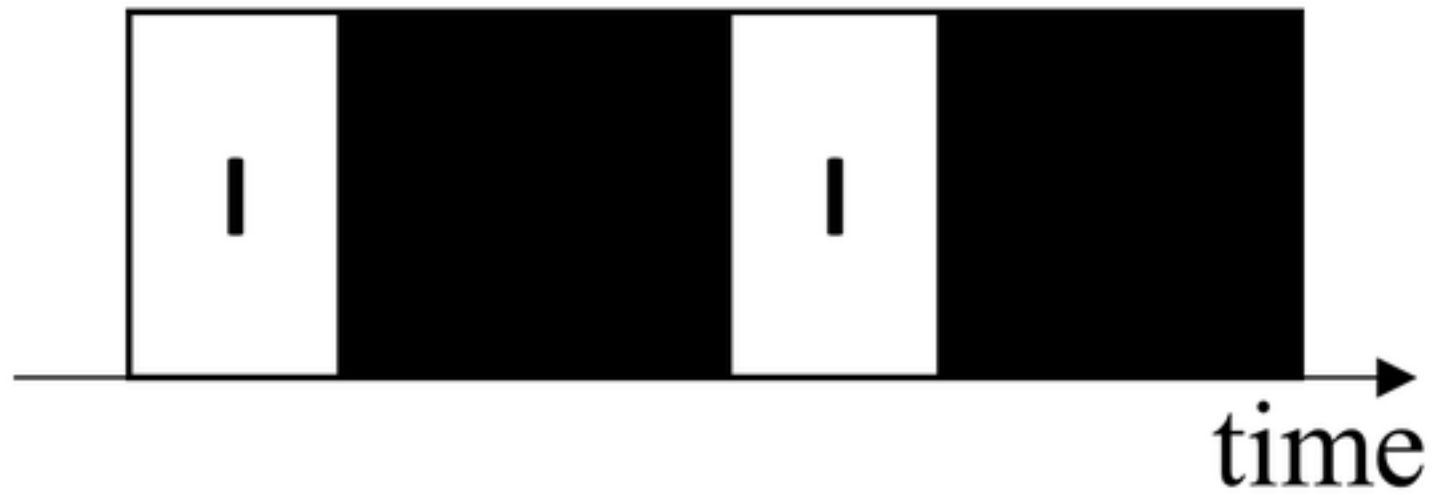
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(a)



(b)



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