

USE CASE

Diagnose-Stratify-Treat: How we designed and deployed predictive AI in a clinical pathway for COPD



CHRIS CARLIN

Consultant Physician & NRS

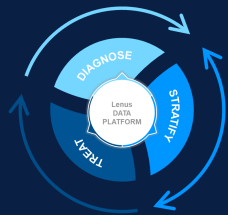
Senior Investigator,

Respiratory Innovation

NHS Greater Glasgow & Clyde



Designing and deploying predictive AI in a COPD pathway



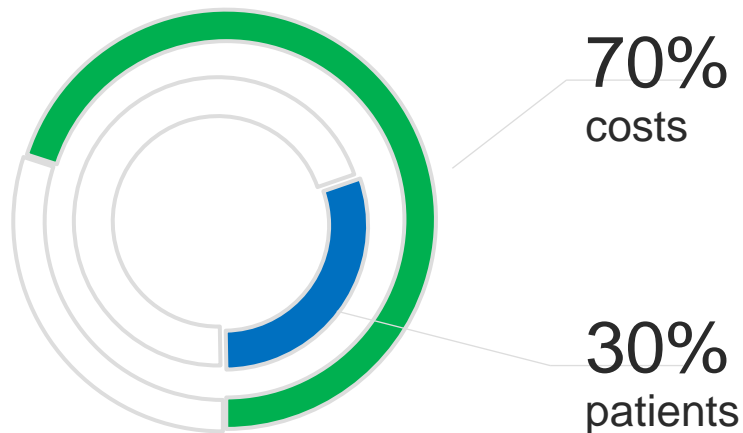
Chris Carlin
Professor, Respiratory Innovation



Lenus

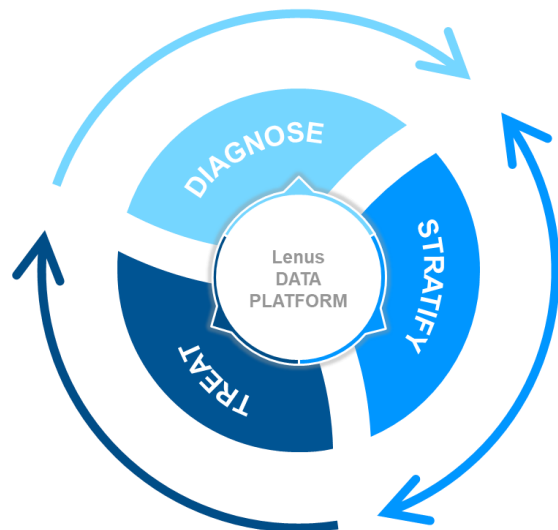
The challenge

Chronic conditions like COPD account for a disproportionate share of healthcare resources and are tied to inequalities



The approach

Build an end-to-end pathway that enables machine learning to support case finding and risk stratification



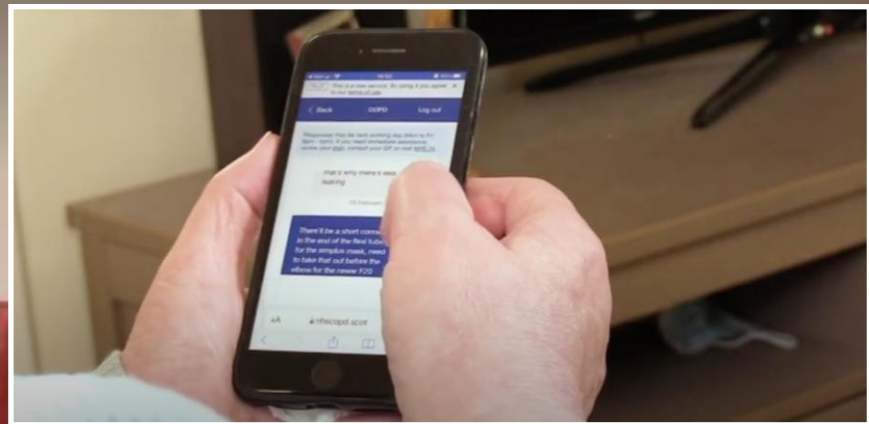
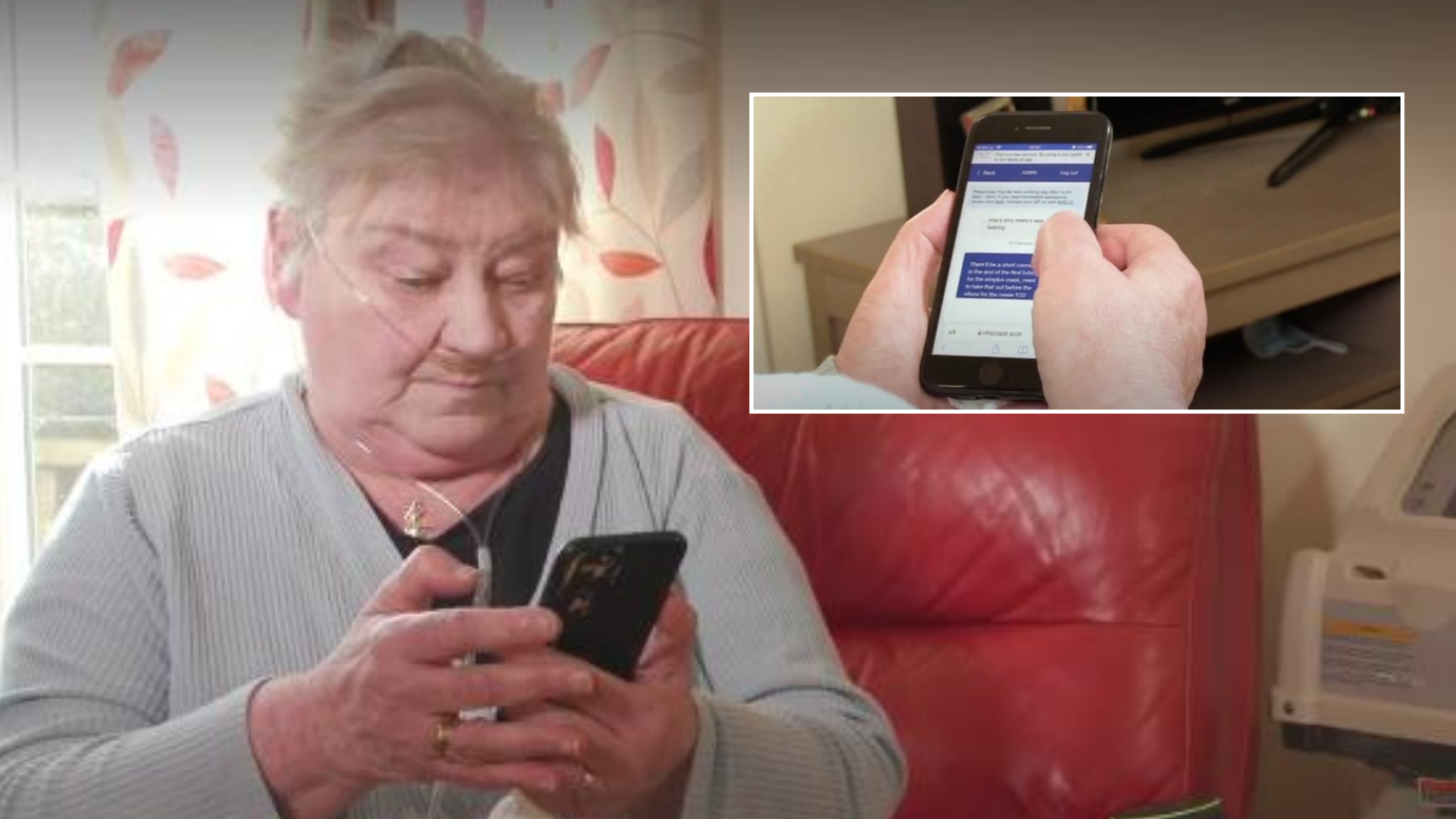


TEST BEDS FOR INNOVATION

Bringing together industry, academia and healthcare services to improve NHS and Social Care in Scotland

GLASGOW Safe Haven

SECURE NHS DATA RESEARCH

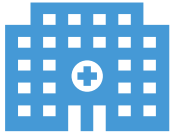




Lenus



1 per week



1 per year



NHS
SCOTLAND
COPD

Joe Sample
Age 62 Sex Male ID 14027250323 Phone no 07700 900000 Next of kin Jere Sample, daughter

Insights

- 72% exacerbation risk (Last updated 8 Dec 2019) 25%
- 12 Month mortality (Last updated 8 Dec 2019) 75%
- 3 Month readmission (Last updated 8 Dec 2019) 42%

Patient reported Outcomes >

- How are you feeling? (Last updated 8 Dec 2019) 4
- CAT (Last updated 8 Dec 2019) 22
- How is your breathing? (Last updated 8 Dec 2019) 2

Last completed

- CAT (04 Aug 2019)
- MRC (04 Aug 2019)
- EQ 5 d (16 Aug 2019)
- Symptom Diary (Never)

Data >

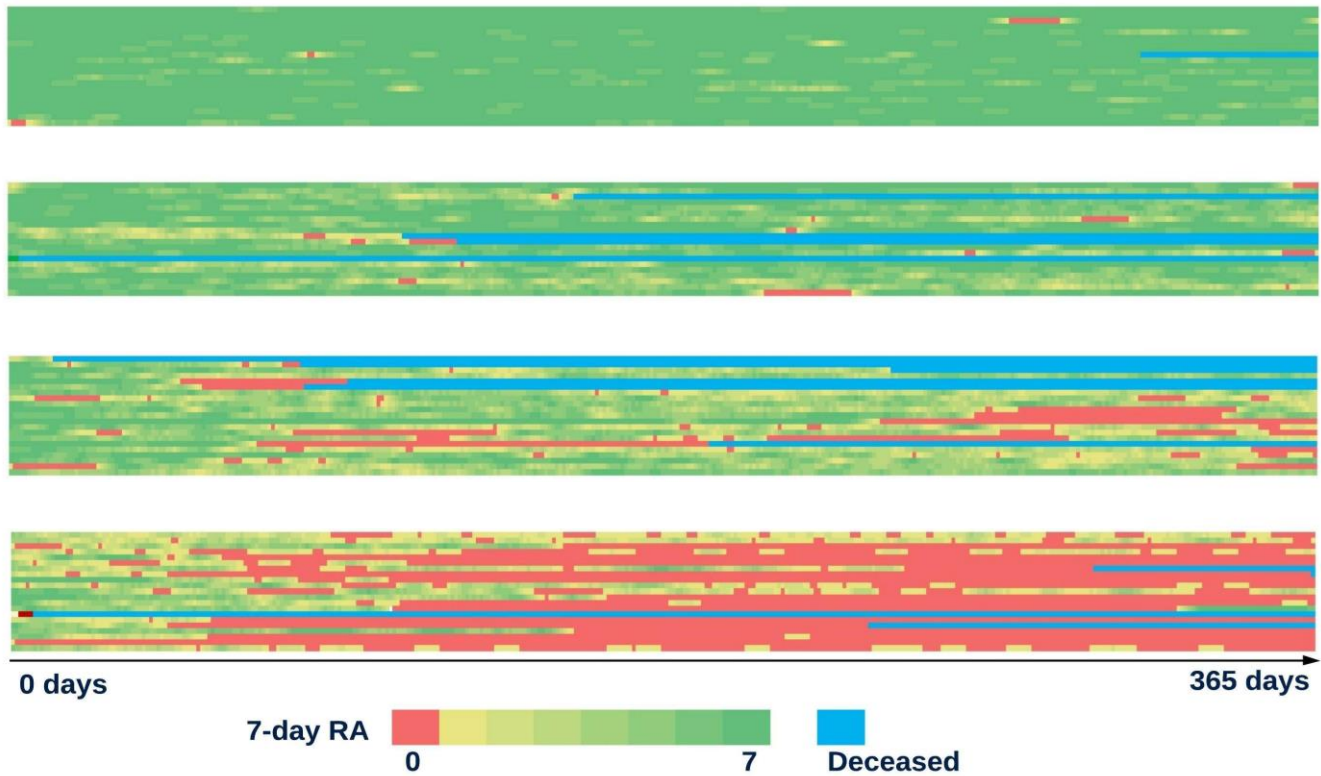
- Steps (Last updated 8 Dec 2019) 2050
- NV Usage (Last updated 8 Dec 2019) 6.5 hrs
- NV Leak (Last updated 8 Dec 2019) 0.5 L/min

COPD Status

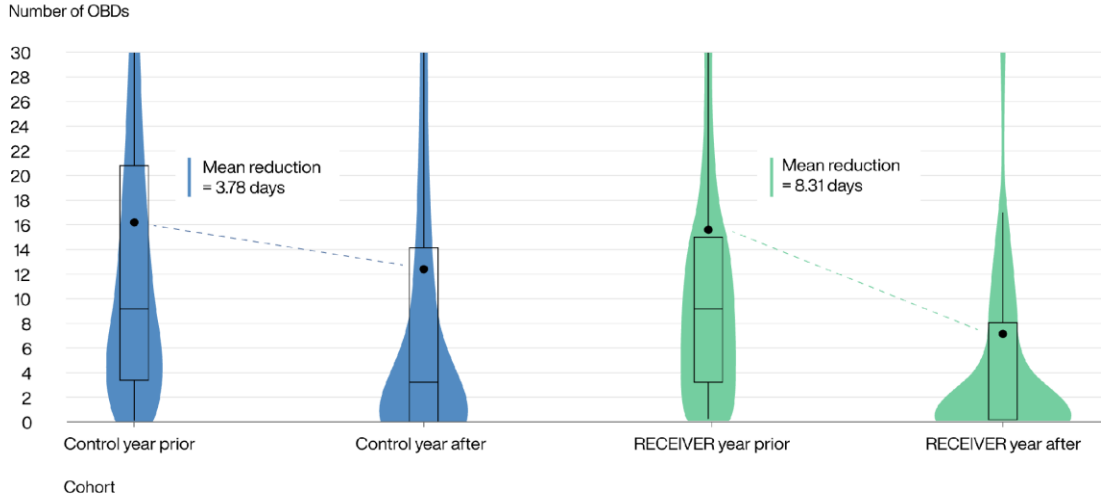
- MRC 4
- FEV1 11
- FEV1 predicted 31%

How are you feeling today?

- Better than usual
- Normal/usual
- Worse than usual
- Much worse than usual



lenushealth.com/evidence



● Control ● RECEIVER

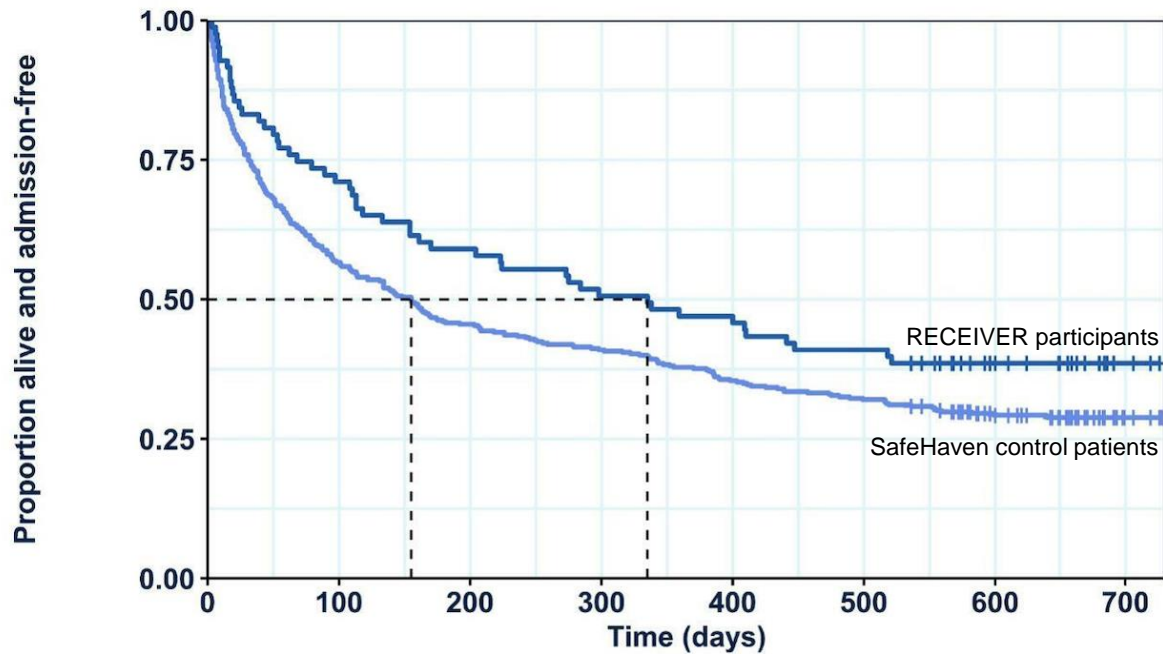
 500 patients

Healthcare resource utilisation saving

£ 3.4m per year



lenushealth.com/evidence



Percentage at risk

	0	100	200	300	400	500	600	700
Control	100	57	46	41	35	32	20	3
RECEIVER	100	71	59	51	47	41	24	4





“ This is just the tip of
the iceberg... ”

Predicting 12-month mortality in a Scottish COPD cohort

De-identified cohort established from NHS GG&C SafeHaven

- NHS GG&C: largest healthcare organisation in UK; 1.2m population, high prevalence of COPD with high admission & mortality rates¹.
- 55531 patients with COPD (ICD-10 J44* in NSS SMR01 dataset)
- Index severe exacerbation + minimum 12 months follow-up data
- Demographic, coded diagnoses, hospital admission, prescribing and laboratory data.

Predictive model co-designed by clinician and data science team

- Binary target variable: alive or deceased at 12 months following severe exacerbation?
- Class stratified 15% of data (7884 patients) = hold out test dataset.
- Remaining 85% of data used for model training and k-fold cross validation.
- ML models applied, with XGBoost demonstrating best performance.
- Model features and performance on hold out test dataset presented.

1. *Scottish Atlas of Healthcare Variation, ISD Scotland.*

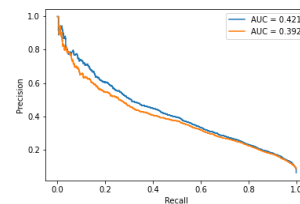
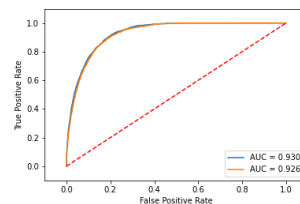
https://bit.ly/COPD_ScotAOV

Shane Burns¹, Grace Cox¹, David J Lowe², Anna Taylor², Paul McGinness¹, Chris Carlin²

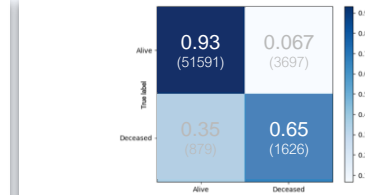
¹ StormID, Edinburgh ² Respiratory & Emergency Medicine, NHS Greater Glasgow & Clyde



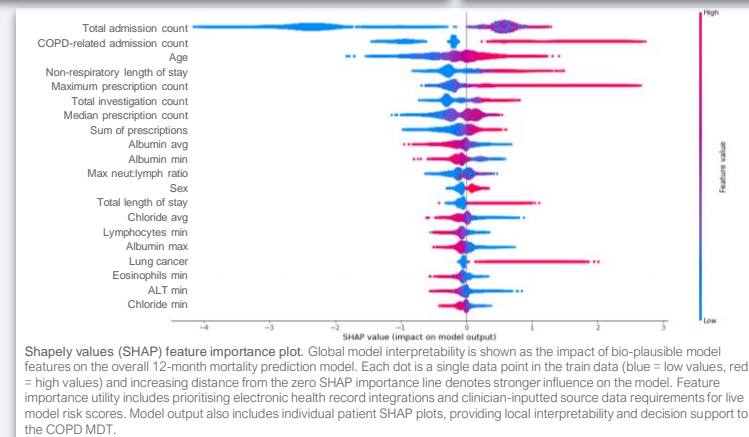
stormid



Receiver operator characteristic and precision recall curves. Red line on ROC curve is a no skill model and blue lines on both curves are baseline model performance on the holdout dataset (reported as the area under curve, AUC). Prediction of alive vs deceased at 12 months following a severe COPD exacerbation exceeds published comparators, with high accuracy and precision supporting clinical implementation. Model performance is retained in dropout analysis (orange lines) when comorbidity feature set is rationalised from all coded diagnoses to a data-driven 'top 20' diagnosis list which can be realistically captured in the LenusCOPD clinical user interface



Model confusion matrix from patient-year observations. Actionable AI insights for an individual can be derived. For example, a patient predicted by this model to be alive has a post-exacerbation 1.7% 12-month mortality. A patient predicted to be deceased has a post-exacerbation 30.5% 12-month mortality, which triggers MDT discussion and prioritisation for anticipatory care planning. 9% of patients fall into this deceased prediction category, which is a manageable MDT workload.



Shapely values (SHAP) feature importance plot. Global model interpretability is shown as the impact of bio-plausible model features on the overall 12-month mortality prediction model. Each dot is a single data point in the train data (blue = low values, red = high values) and increasing distance from the zero SHAP importance line denotes stronger influence on the model. Feature importance utility includes prioritising electronic health record integrations and clinician-inputted source data requirements for live model risk scores. Model output also includes individual patient SHAP plots, providing local interpretability and decision support to the COPD MDT.

Our AI-based COPD 12-month mortality prediction model demonstrates excellent performance: integration of model scores and explainability plots within LenusCOPD "AI Insights" dashboard extension for MDT use is planned.

Experience with MDT presentation of model risk scores in our proposed 'DYNAMIC-AI' implementation-effectiveness clinical investigation will inform adoption and further evaluations of AI insight-based decision support.

Additional models - 3-month admission and 72-hour exacerbation risk - are in advanced development.

Daily patient-reported outcome with wearable and respiratory therapy monitoring data from scale-up of the LenusCOPD service (support.nhscopd.scot) will provide reference ground truth event data and input features for continuous model improvements.

Predicting 3-month respiratory readmission in a Scottish COPD cohort

De-identified cohort established from NHS GG&C SafeHaven

NHS GG&C Largest healthcare organisation in UK, serving 1.2m population. High prevalence of COPD, with high admission & mortality rates.

Dataset Demographics, coded diagnoses, hospital admissions, prescribing and laboratory data from 33148 patients with COPD who had 120252 respiratory-related admissions.

Predictive AI model co-designed by clinician and data science team

Binary target variable Respiratory-related readmission at 3 months post discharge.

Model training 85% of data used, with k-fold cross validation.

Holdout test dataset Class stratified 15% of data to evaluate models' performance and fairness.

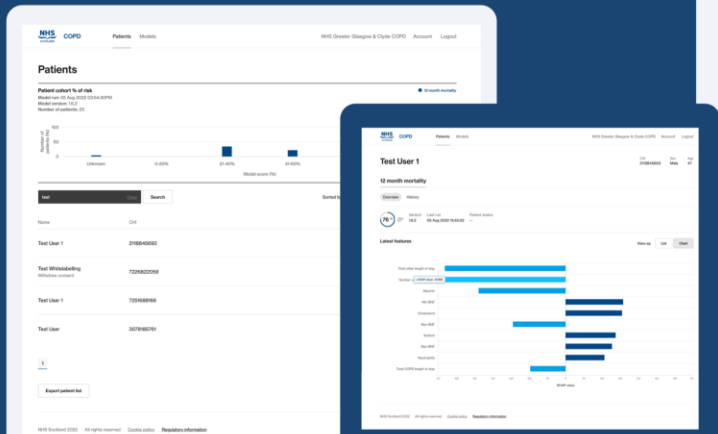
XGBoost model Selected as demonstrated best performance.

Conclusions

Model's performance and utility Ready for adoption within COPD MDTs.

Operationalising live AI models Co-designed cloud-based COPD AI insights app ready to be deployed.

DYNAMIC-AI Patient acceptability and technical feasibility will be determined alongside a range of safety and utility secondary objectives in this implementation-effectiveness trial.

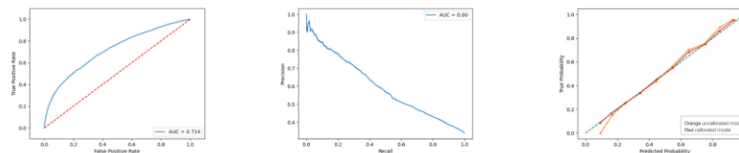


Shane Burns, Anna Taylor*, Grace Subašić, Paul McGinness, David J Lowe*, Chris Carlin*

Lenus-Health, Edinburgh and *Respiratory & Emergency Medicine, NHS Greater Glasgow & Clyde

Get in touch lenushealth.com or christopher.carlin@ggc.scot.nhs.uk

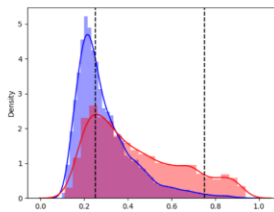
Receiver operator characteristic, precision recall and calibration curves from holdout test dataset



Distribution of model inference probabilities

Blue no readmission Red readmission

Selecting probability threshold of 0.25 tunes model to bring forward most patients with readmission, with reduced specificity. Selecting probability threshold of 0.75 tunes model to high specificity, but misses large n of patients who have a readmission. Clinicians & data scientists can collaborate to adapt model outputs depending on use case and service capacity.

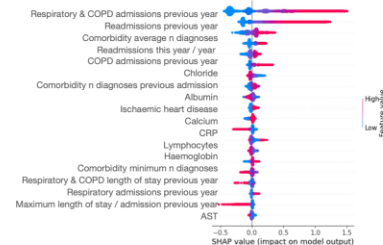


Probability threshold	Number of patients predicted 3-month readmission / 100 patients		
	Correct	Incorrect	Missed
0.25	27	34	7
0.5	12	5	23
0.75	4	0	31

Global model explainability data

SHAP plot of model feature importance

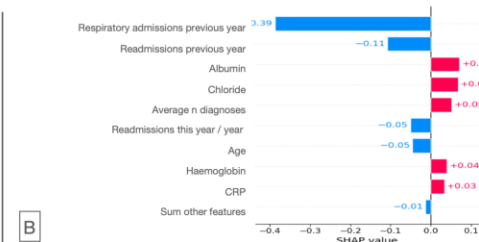
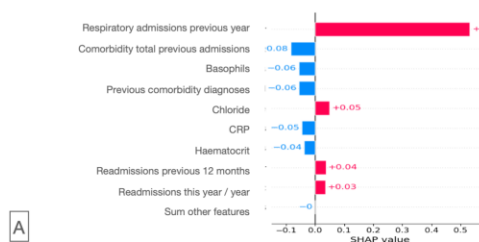
Feature review provides actionable insights: respiratory failure, requirement for respiratory support and anxiety are highlighted as diagnoses associated with high-risk for readmission.



Individual patient explainability data

SHAP plots of model feature importance

A Correct prediction readmission B Correct prediction no readmission



Model development dataset

- NHS Greater Glasgow & Clyde Safe Haven DYNAMIC cohort.
- All patients with COPD resident NHS GG&C 1st Jan 2013 - 31st Dec 2019.
- Datasets: demographics, prescribing, laboratory, hospital admission and comorbidity data.

Changes from previous model:

- None

Model formulation

- Binary classification problem predicting 12-month mortality (Class 0 = alive at 12 months; Class 1 = deceased at 12 months).
- Model inference per patient per model run yields a score ranging from 0-1, which can be interpreted as the probability of patient being deceased at 12 months. This probability is converted to a percentage in the COPD AI Insights App.
- XGBoost algorithm (decision tree based) with hyperparameters tuned to the clinical problem.
- All model parameters and metrics are logged in the model tracking system.

Changes from previous model: None

Model training and validation

85% of dataset (~ 39K patients)

Model evaluation

15% of dataset – 'holdout' test (~ 7K patients)

- In order to prevent data leakage, the full dataset was split such that an individual patient may only appear in either the train or holdout test cohort.
- Demographic make-up (mean age = 68 yr, 57% female) and class balance (proportion of patients alive vs deceased at 12 months, 7%) matched for training and holdout test datasets.
- In order to prevent data leakage, model validation during training was performed using K-fold cross validation, in which the train dataset is split patient-wise into K folds.
- Model performance and associated metrics for model approval and presentation in the COPD AI Insights App are reported from the holdout test cohort.

Changes to this approach from previous model: None

Feature engineering

- Features are a combination of one-hot encoding, target encoding and domain-driven feature engineering techniques. See Table 5 for the model features and applied engineering techniques
- Where target encoding or other data aggregations are performed across patients, this uses a K-fold approach to prevent data leakage. Aggregations from the training data were applied to holdout test data and will be applied to inference data.
- Data quality issues and missingness were explored and remedied where necessary.
 - A maximum missingness threshold of 40% was applied to lab data to determine whether a lab test feature could be included in the model.
 - A small amount of poor-quality pharmacy data was removed.
 - Missing data for the selected laboratory-derived features (those with missingness less than 40%) was allowed during model training. This was the case as the chosen algorithm was sparsity aware, meaning the missing data did not need to be imputed.
 - Missing data in all other features was filled appropriately. For all other features, missing data were replaced with zero as the context for missingness in those cases implied zero occurrence.
- Patient age was scaled using the training data. The holdout test dataset was inflated and scaled using properties of the training data to prevent data leakage.
- Multiple admissions corresponding to the same hospital admission event were consolidated as one admission. This occurs, for example, when a patient is transferred between different hospital facilities as part of the same admission.
- Feature interaction constraints are imposed on the 'days since' lab test features. Each 'days since' feature can only interact with the corresponding lab test, the patient's age, and the patient's sex.
- Hospital admission diagnosis codes are collated to align to comorbidities as recorded in [Lexus COPD](#) service.

Changes from previous model: None

Model performance metrics

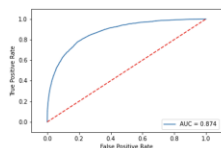


Figure 3 - Receiver Operating Characteristic (ROC) curve of the candidate model, evaluated on the holdout test cohort.

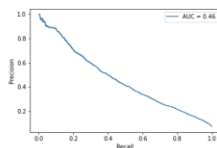


Figure 4 - Precision-Recall (PR) curve of the candidate model, evaluated on the holdout test cohort.

Probability threshold	Class	Precision	Recall	F1 Score	ROC-AUC	PR-AUC	Brier Loss
0.25	Deceased	0.46	0.44	0.45	0.87	0.46	0.048
	Alive	0.96	0.96	0.96			
0.5	Deceased	0.70	0.2	0.32	0.87	0.46	0.048
	Alive	0.99	0.94	0.97			
0.6	Deceased	0.78	0.16	0.26	0.87	0.46	0.048
	Alive	1	0.94	0.97			
0.8	Deceased	0.88	0.08	0.15	0.87	0.46	0.048
	Alive	1	0.94	0.97			

Table 2 - Candidate model performance with different probability thresholds, evaluated on the holdout test cohort.

Comments:

- The candidate model performance is in line with expectations.
- Candidate model performance is the same as the current model.

Model calibration, performance and thresholding

- If a model is perfectly calibrated, the inference probabilities will match the true probability of the event, e.g., half of patients with a model score of 0.5 will truly be deceased within 12 months.
- The candidate model was calibrated during cross validation to maximise matching between inference probability and true probability.
- Thresholding is the process of converting probability outputs to class labels by choosing a cut off value (values less than this output as class 0, values greater than this as class1). The choice of threshold will affect the model performance metrics and expected clinical workload
- Several model performance metrics are presented in this report. Of particular importance for this clinical use-case are:
 - The area under the precision recall curve (PR-AUC). Other metrics tend to be inflated and misleading when class imbalance is high.
 - Expected numbers of patients brought forward correctly/incorrectly and missed for different threshold levels. This is important as it simulates the expected clinician workload.

Changes to this approach from previous model:

- Updated calibration method to use custom patient folds versus the built in [sklearn](#) record-wise fold generation. This gave a slight improvement to model calibration.

Model explainability and fairness

- In accordance with ethical AI principles, model explainability and fairness were evaluated
- Global model explainability describes which model features are important to the overall model across the entire training cohort.
- Local model explainability describes what is important on an individual prediction level.
- This allows for interrogation of the model prediction and to identify potential biases.
- Model fairness relates to performance on different sub-groups of interest within the population.
- These sub-groups may relate to demographics (e.g. sex, age, ethnicity) or clinical factors such as the presence/absence of certain comorbidities.
- The model can be re-trained if potential inequalities are identified, tuning the loss functions to ensure parity between groups.

Changes to this approach from previous model:

- The [shap](#) values for both global and local explainability are now calculated by averaging the [shap](#) values from each of the base models that form the final calibrated model.

Model QA (technical)

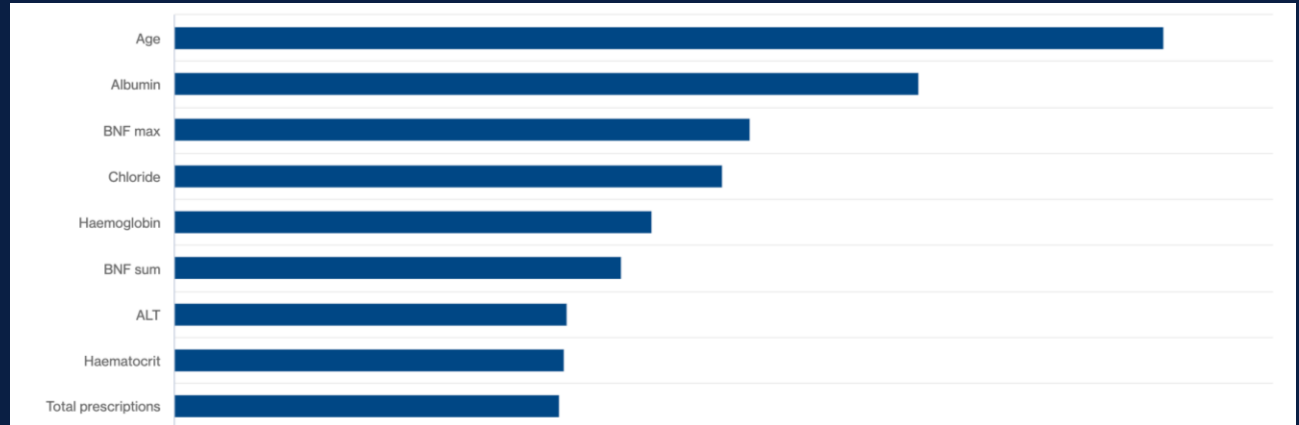
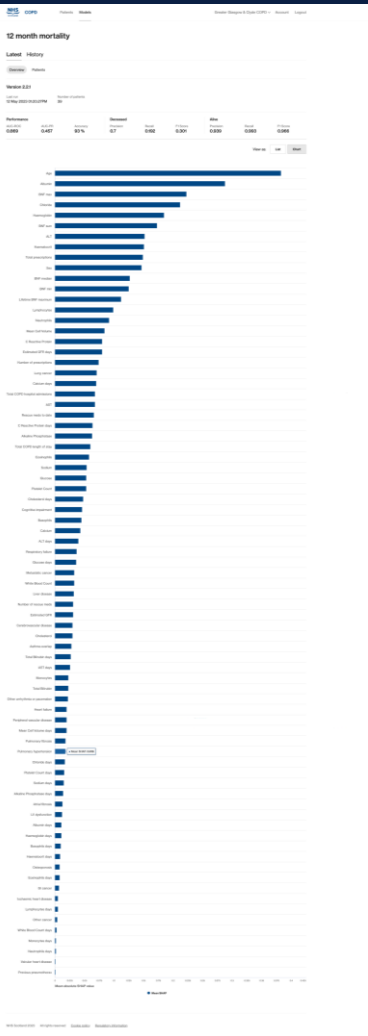
The model was reviewed internally by the [Lexus](#) Engineering team on 31/10/2022 and deemed an acceptable candidate for clinician model approval. The [detailed report](#) of the technical review is available.

Model approval review

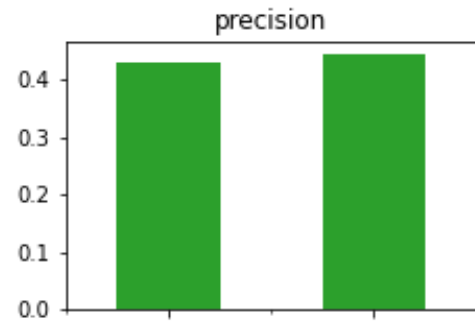
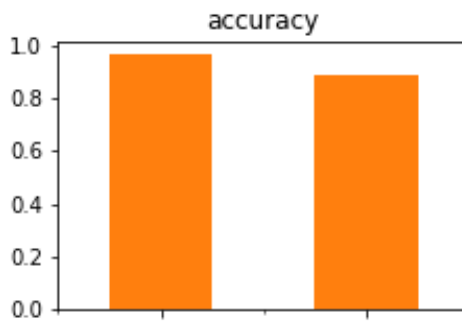
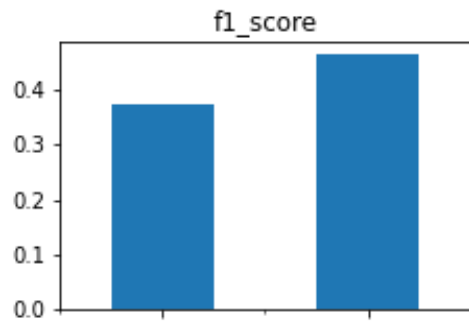
Highlights and discussion points

- The candidate model is well calibrated. This can be seen visually from the calibration curve, and from the low Brier Loss.
- There is reasonable separation of inference probabilities for the two classes.
- A clinical decision based on available resources is required for the most appropriate threshold.
- Model performance is in line with expectations and unchanged from the current model.
- Checks in cross-validation indicate that no data leakage or overfitting to training data has occurred.
- The candidate model global and local explainability appears [bio-plausible](#). To be verified with clinicians.
- The candidate model performs slightly better on men versus women.
- The candidate model performs better on the over 65s age group vs under 65s.
- The candidate model performs slightly better on less deprived SIMD groups.
- The context of having had a specific lab test can be more important than the value itself. There are more missing values for people who did survive the following year implying that people who are generally sicker, get tested for more things. This motivated the decision to impose interaction constraints on the lab 'days since' variables to help improve model generalisability. In a different healthcare setting it may be more routine to take certain labs test measurements so using the 'days since' feature without interaction constraints may introduce a systematic bias.

Model explainability



Model fairness



Dynamic AI study

We're running a study to find out if computer-based problem solving, or artificial intelligence (AI), can help us care for patients with COPD.

About the study

Find out more about the study, including how it works and who's running it.

Your privacy

How we're keeping your information safe, secure, and confidential.

Join the Dynamic-AI study

Dynamic-AI is available by invitation only for people with COPD in Greater Glasgow and Clyde.

The screenshot shows the website's navigation bar with links for Home, Managing my COPD, About this service, Setup, Dynamic AI, and Join. The main content area includes:

- About the study:** A blue header with the text "Find out more about the study, including how it works and who's running it."
- About Dynamic-AI:** A section with a video player titled "What is the Dynamic AI project?". The text above the video states: "We're investigating whether we can use artificial intelligence (AI) to improve the healthcare of people with COPD."
- Why AI?** A section with a video player titled "Will AI be replacing the doctors?". The text above the video states: "Through the COPD app and electronic patient records, we collect a lot of information. Analysing this information takes a lot of time. AI is a set of computer programs that can quickly and accurately analyse this information and bring out highlights. Doctors will then use these highlights to make decisions about patient care. AI will not replace any of your existing care; doctors will simply use it to help them make decisions. This way can spend less time looking through all of the information and more time with patients."
- How the AI works:** A section with a video player titled "Will AI be replacing the doctors?". The text above the video states: "The Dynamic-AI model is a computer program that looks for patterns in health information. The AI model would use information from both the COPD service and your electronic health record. For example, through factors in your app, journal, and any medication you might be using. The AI model aims to use this information to assess our your likelihood of a change or deterioration in your COPD. This information is gathered and would be used to give a broader picture of your individual health. The goal is to find out both when your health is at risk, and why."



DYNAMIC-AI
Support website

COPD AI insights

Find out more about this study



Questions

It's time to answer your COPD questions

Start

Self management



Messages



Settings and privacy



[Get help from the support site](#)

COPD AI Insights

What we need from you

When you join the study, we will access information from:

- your daily COPD questions
- your Fitbit, if you have one
- your ResMed kit, if you have one

We'll use this information to run and improve our AI systems.

We will record this information in your electronic health records.

Next

COPD AI Insights

Your privacy

Your identity and your personal information will be secure and kept private. All the information we collect as part of this study will be stored electronically by NHS GG&C.

Your information may be accessed by:

- NHS GG&C research team
- Commercial partners at Lenus Health and Storm ID
- Regulatory bodies making sure the study is being run correctly

Only the clinical team at NHS GG&C will be able to see your identity. Commercial partners and regulatory bodies will not be able to access your personal information.

COPD AI Insights

Consent

Before you join the study, we need your permission to access and use your information.

Model operation

We need to use your information to:

- use the artificial intelligence model
- help doctors make decisions
- help to make decisions about patient care

I consent

Model training

We need to use your information to:

COPD Insights

Sign in to the COPD Insights service.

[Log In](#)

Patients

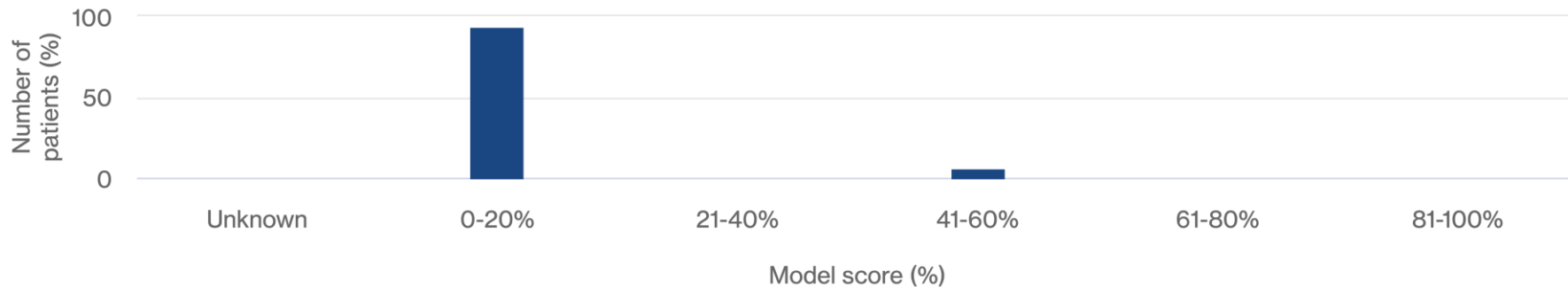
Patient cohort % of risk

● 12 month mortality

Model run: 19 Apr 2023 04:21:23PM

Model version: 2.2.1

Number of patients: 17



12 month mortality

Overview History

57%

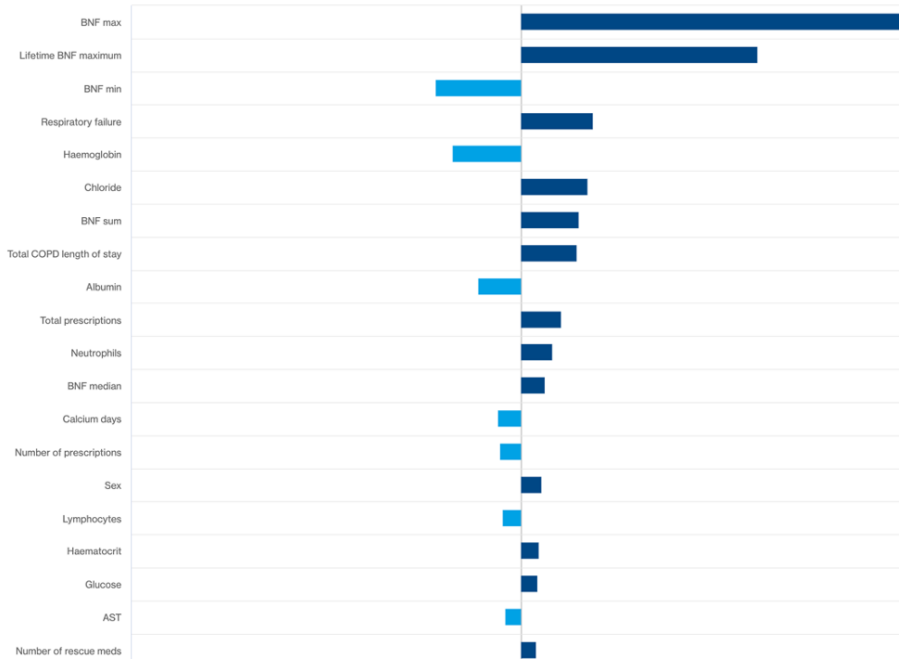
Version 2.2.1 Last run 19 Apr 2023 04:21:26PM Patient status --

Latest features

View as:

List

Chart



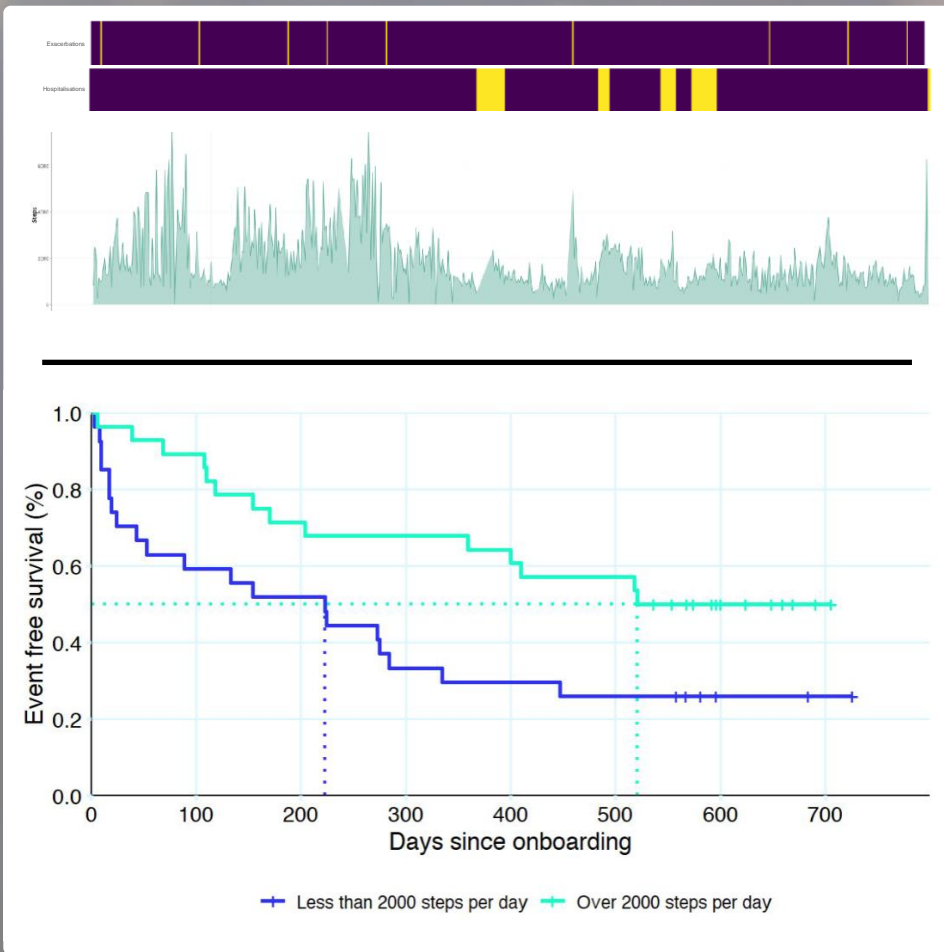
Hi Mr

Thanks for joining the DYNAMIC-AI trial. We got the first run of data through from it, and it's really interesting, with nothing worrying. It did flag one thing - that it might be worth checking your overnight breathing or blood gases again at some point. We could have a chat about that - not urgent, but I'd have clinic space + could give you a call tomorrow or thursday sometime, if any time either day would suit you? Chris

Chris Carlin - 25 April 2023 13:28



“ This is just the tip of
the iceberg... ”





12-month admission vs no admission

Parameter	Mean- No Admissions	Mean- Had admissions	SD- No admissions	SD- Had admission	p value (Mann-Whitney test)	Hodges Lehmann estimate	Hodges Lehmann estimate/ Mean- No admissions
IE ratio (Max)	64.189	58.150	19.809	16.224	6.71e-37	5.9999795	0.0934736
IE ratio (Median)	40.133	34.819	10.053	7.015	1.04e-111	5.9999555	0.1495018
IE ratio (95th)	54.056	45.064	14.158	10.525	1.55e-154	9.0000309	0.1664946
Minute ventilation (Median)	9.341	8.876	2.106	2.243	3.84e-53	1.1250278	0.1204398
Minute ventilation (95th)	12.107	11.517	3.277	3.881	7.73e-36	1.2500347	0.1032489
Apnea-hypopnea index	1.855	1.114	3.242	2.898	1.14e-109	0.5000382	0.2695623
Hypopnea index	1.505	0.606	2.948	1.263	5.80e-129	0.4000876	0.2658390
Patient triggered expiration (%)	67.994	84.763	22.033	20.435	2.26e-272	-16.4999548	-0.2426678
Patient triggered inspiration (%)	76.343	81.758	21.375	24.747	7.00e-67	-5.0000172	-0.0654941
EPAP value (Median)	7.669	9.019	1.734	2.919	8.22e-50	-1.1999474	-0.1564673

90-day admission vs no admission

Parameter	Mean- 90 days pre-admission	Mean- Other	SD- 90 days pre-admission	SD- Other	p value (Mann-Whitney test)	Hodges Lehmann estimate	Hodges Lehmann estimate/ Mean- 90 day pre-admission
Leak value (Max)	0.927	0.622	0.693	0.675	5.37e-27	0.2800298	0.3020818
Leak value (95th)	0.460	0.244	0.479	0.378	1.12e-31	0.1399548	0.3042496
Minute ventilation (95th)	13.358	10.919	4.999	3.222	2.43e-20	1.7500286	0.1310098
Apnea-hypopnea index	2.866	0.545	5.049	1.236	1.69e-46	0.3999887	0.1395634
Apnea index	1.457	0.178	3.350	0.722	3.02e-48	0.0999804	0.0686207
Hypopnea index	1.384	0.353	2.041	0.706	1.80e-44	0.2000183	0.1445219
Respiratory rate (Max)	30.129	25.042	6.067	4.523	4.57e-89	5.6000694	0.1858697
Respiratory rate (Median)	19.436	16.695	4.875	2.949	6.00e-34	2.2000544	0.1131948
Respiratory rate (95th)	25.253	20.555	6.166	4.243	3.32e-74	4.7999485	0.1900744
Minutes spO2 below 88 percent	2.502	1.994	1.026	0.850	1.84e-33	0.4805169	0.1920531
Seconds spO2 below dynamic threshold	0.056	0.050	0.020	0.021	6.83e-13	0.0060204	0.1075064
EPAP value (Median)	9.803	8.764	2.691	2.946	3.17e-24	0.9599499	0.0979241

7-day admission vs no admission

Parameter	Mean- week pre-admission	Mean- Other	SD- week pre-admission	SD- Other	p value (Mann-Whitney test)	Hodges Lehmann estimate	Hodges Lehmann estimate/ Mean- week pre-admission
Respiratory rate (Max)	26.189	29.550	5.351	6.225	2.81e-07	-3.399929	-0.1150568
Respiratory rate (95th)	21.616	24.583	5.128	6.421	7.98e-06	-2.999963	-0.1220340

RECEIVER trial, unpublished data

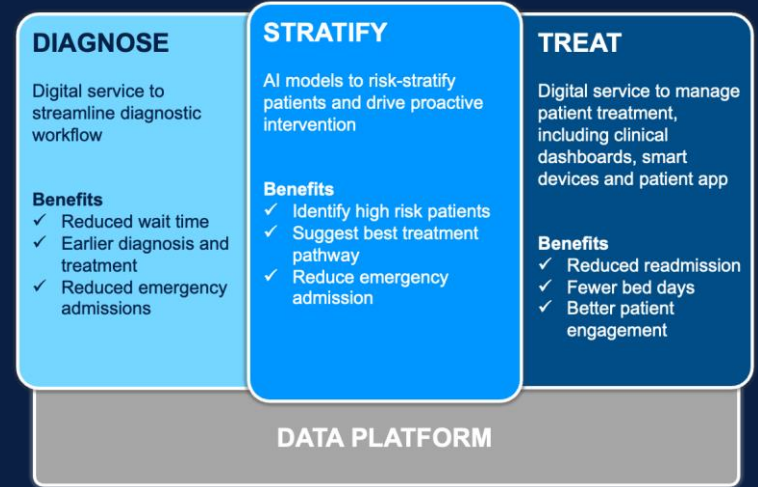
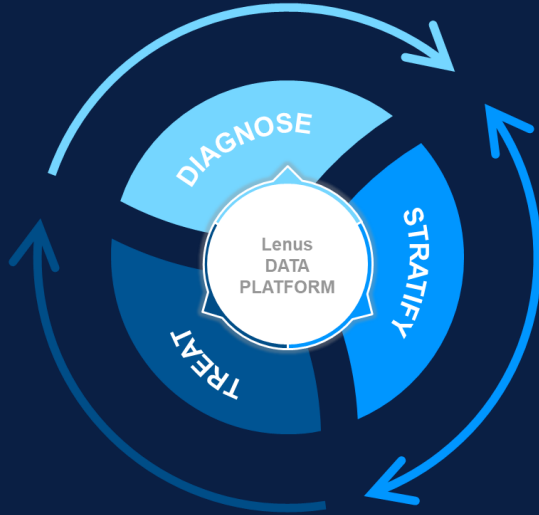


POLARIS

Prevention of COPD Lung Attacks
by Early Intervention Strategies

Predictive AI in an end-end COPD pathway

Exemplar for long term condition care



DYNAMIC-AI trial

Co-design live AI insights at point of care in COPD MDT

Clinical investigation acceptability, feasibility and utility

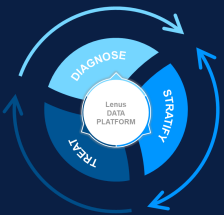
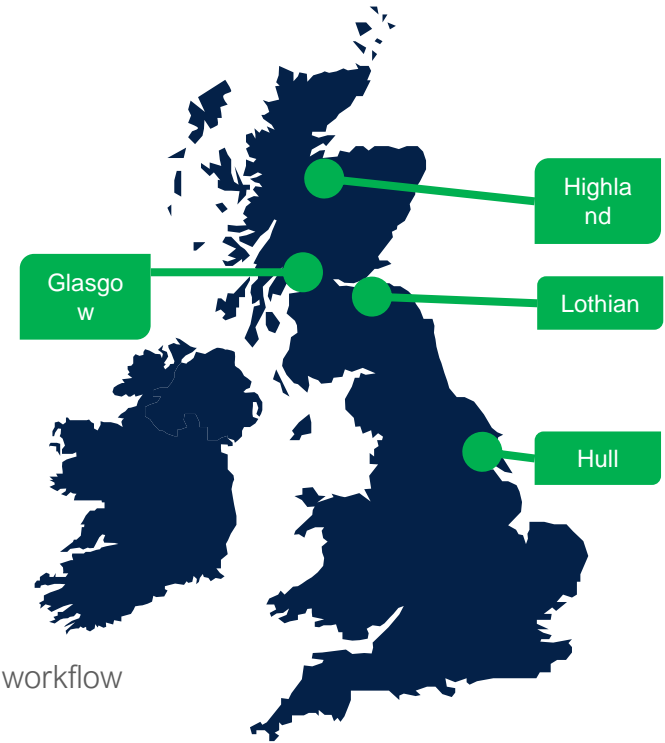
End-end pathway

LenusCOPD service sustained use, improved outcomes

Scale-up recover and re-orientate diagnostics, other sites

Test bed

Implement and validate AI models and other solutions within clinical workflow



Designing and deploying predictive AI in a COPD pathway



Chris Carlin
Professor, Respiratory Innovation



Lenus

INTELLIGENT HEALTH UK 2023

Breaking down the barriers
between tech and healthcare