

## Abstract

**Background:** Life-saving emergency major resection of colorectal cancer (CRC) is a high-risk procedure. Accurate prediction of post-operative mortality for patients undergoing this procedure is essential for both healthcare performance monitoring and pre-operative risk assessment. Risk-adjustment models for CRC patients often include patient and tumour characteristics, widely available in cancer registries and audits. We investigated to what extent inclusion of additional physiological and surgical measures, available through linkage or additional data collection, improves accuracy of risk models.

**Methods:** Linked, routinely-collected data on patients undergoing emergency CRC surgery in England between December 2016 and November 2019 were used to develop a risk model for 90-day mortality. Backwards selection identified a 'selected model' of physiological and surgical measures in addition to patient and tumour characteristics. Model performance was assessed compared to a 'basic model' including only patient and tumour characteristics. Missing data was multiply imputed.

**Results:** 846 of 10,578 (8.0%) patients died within 90 days of surgery. The selected model included seven pre-operative physiological and surgical measures (pulse rate, systolic blood pressure, breathlessness, sodium, urea, albumin, and predicted peritoneal soiling), in addition to the ten patient and tumour characteristics in the basic model (calendar year of surgery, age, sex, ASA grade, TNM T stage, TNM N stage, TNM M stage, cancer site, number of comorbidities, emergency admission). The selected model had considerably better discrimination compared to the basic model (C-statistic: 0.824 versus 0.783, respectively).

**Conclusion:** Linkage of disease-specific and treatment-specific datasets allowed the inclusion of physiological and surgical measures in a risk model alongside patient and tumour characteristics which improves the accuracy of the prediction of the mortality risk for CRC patients having emergency surgery. This improvement will allow more accurate performance monitoring of healthcare providers and enhance clinical care planning.

## Keywords

colorectal cancer; emergency surgery; risk model; post-operative mortality; record linkage; electronic health records.

## Introduction

Major resection is a common treatment for patients diagnosed with colorectal cancer (CRC), and is associated with a high risk of death when undertaken in the emergency setting.[1] Comparisons of post-operative mortality among hospitals, teams or surgeons, or over time, are important for quality assessment and quality improvement of CRC services and risk-adjustment is needed to ensure fair comparisons. Risk models are also important for pre-operative risk assessment, which can aid clinical care planning and inform the counselling of patients for emergency CRC surgery. Clinical guidelines recommend that all high-risk surgical patients should receive certain standards of care, such as direct transfer to critical care and presence of a consultant surgeon and anaesthetist in theatre.[2] Accurate risk prediction is therefore vital to ensure the highest risk patients receive appropriate care.

There are four major arguments to develop a new extended risk model for emergency CRC surgery. The first argument is that recent reviews have identified a substantial number of models for CRC surgery,[3] and for emergency bowel surgery,[4,5] but only a few models for the intersection between these two groups. Using either a CRC risk model or an emergency bowel surgery risk model may lead to inaccurate predictions of risk for patients undergoing emergency CRC surgery.

Second, the available models that focus on emergency CRC surgery were specifically developed using data from a single provider,[6,7] or from a limited geographical area.[8] A risk prediction model developed in a national population of patients undergoing emergency CRC surgery will have increased precision in the model estimates and will be more widely applicable.

Third, risk models for CRC surgery tend to include patient and tumour characteristics, which are widely available in cancer registries and audits of care for patients with CRC. For this new risk model, we also considered the inclusion of physiological measures (e.g., measurements of organ function and overall health, such as serum creatinine level, breathlessness history, etc.) and surgical measures (e.g., peritoneal soiling, intraoperative blood loss, etc.), which are less readily available. Their inclusion may improve accuracy of risk prediction in the emergency setting.

Fourth, increasing the number of measures included in a risk model may also limit the model's utility in clinical practice given the potentially greater impact of missing data, mismeasurement, and misclassification. We therefore used an explicit model development approach to ensure that the improvement in prediction accuracy by including physiological and surgical measures is balanced against utility of the model.

The development of such an extended risk model for patients undergoing major emergency CRC resection is possible due to availability of detailed patient, tumour, surgical, and physiological information from a large cohort of patients recorded in linked electronic health databases.[9] In the study described in this paper, we investigated whether accuracy of a basic risk model including only patient and tumour characteristics was improved by inclusion of physiological and surgical measures. We aimed to develop and validate such an extended risk model that can be used for both *risk-adjustment* for performance monitoring of healthcare providers, and *risk prediction* for clinical care planning.

## Methods

### Datasets and linkage

Data from disease, treatment and administrative hospital databases for patients in England were linked in a national cohort study. For information on patient characteristics, hospital admissions and outcomes, we used an administrative hospital database, Hospital Episode Statistics Admitted Patient Care (HES), which contains information on all hospital admissions in the English National Health Service (NHS).[10,11] For information on patient and tumour characteristics, we used a disease-specific dataset collected by the National Bowel Cancer Audit (NBOCA), containing data on patients diagnosed with CRC.[1] For physiological and surgical measures, we used a treatment-specific dataset collected by the National Emergency Laparotomy Audit (NELA), containing data on patients having emergency bowel surgery.[12] Each of these datasets contains information on mortality provided via national UK mortality statistics.[13]

The three datasets were linked using a spine linkage approach, where the administrative hospital database was designated the 'spine dataset' and the other two datasets were linked to it.[9] Linkage used deterministic rules regarding agreement on patient identifiers (NHS number, sex, date of birth, and residential post code).[9,14]

Patients were eligible for analysis if they were in the spine dataset, were recorded as undergoing an emergency major resection for CRC between 1 December 2016 and 30 November 2019 in at least one dataset (Appendix: Table A.1, Figure A.1), and had complete data for mortality.

The eligible cohort was split into development and validation datasets based on date of surgery. The development cohort included patients having surgery between 1 December 2016 and 30 November 2018. The validation cohort included patients having surgery between 1 December 2018 and 30 November 2019.

## Identification and definition of variables

The risk model for 90-day mortality was developed to be used for both risk-adjustment and for clinical care planning. For both risk-adjustment and clinical care planning, variables need to capture a patient's risk immediately *before* surgery. For risk-adjustment, variables included need to be factors outside of the control of the provider, so that due merit is given to providers with high-quality care. For clinical care planning, information that is reflected by the included variables needs to be available before surgery. Variables to be considered for inclusion were identified from existing risk models for CRC surgery or emergency bowel surgery.[3-5,15-21]

A 'basic model' included *all* patient and tumour characteristics included in these models that met the inclusion criteria and were routinely available in electronic health records (i.e., without any variable selection). Physiological and surgical measures included in models identified in the literature were candidate variables for selection if they met the inclusion criteria, and were available through linkage to external databases. Candidate surgical measures were pre-operative estimates of operative severity (i.e., type of procedure to be undertaken), expected peritoneal soiling, and expected intraoperative blood loss, as these variables reflect clinicians' assessments of patients' health just before surgery.[16]

For continuous variables, we used fractional polynomials with functional forms established in an existing model developed and validated in 38,830 patients (e.g. age was modelled as linear plus quadratic).[16] To reduce the influence of outliers, continuous variables were Winsorised, by setting observed values beyond the 1st and/or the 99th percentile as the value of that percentile (Appendix Table A.2). Levels of categorical variables were combined if any categories had <10 patients with complete data in either the development dataset or validation dataset. We considered all interactions that had been identified by existing models: an interaction between age and metastases, an interaction between age and ASA grade, and an interaction between ASA grade and respiratory history.[15,16]

## Handling missing data

Multiple imputation with chained equations was used to handle missing data, under the assumption of missing-at-random conditional on mortality and other observed information.[22] The imputation procedure was undertaken separately in the development and validation datasets, and for the full dataset as a whole. The number of imputations was set at 20. Year of procedure, sex, and procedure

type were complete for all patients. 8 patients with missing mortality were excluded from analysis (Appendix Figure A.1). Binary variables were imputed using logistic regression, categorical variables were imputed using multinomial logistic regression, continuous variables were imputed using predictive mean matching with values drawn at random from a pool of k=3 observations with similar predicted values.[23,24]

Parameters and performance measures (on the appropriate scale) were pooled over imputed datasets using Rubin's rules.[25] As the  $\chi^2$  statistic from Hosmer-Lemeshow (HL) tests cannot be pooled using Rubin's rules, we calculated a F-statistic to account for between-imputation variation.[26]

Potential bias from non-linkage, the main source of missing data, was assessed by comparing patient characteristics (recorded in the administrative spine dataset) and 90-day mortality between linked and unlinked patients.[27]

## Model development

Using the development dataset, a 'full model' was fitted including all patient and tumour characteristics in the basic model and all candidate physiological and surgical measures. To assess whether a simpler model could achieve similarly high prediction performance, backwards selection was used with criteria defined by model  $R^2$  (measure of variation in outcomes explained). At each step,  $R^2$  values were pooled over the imputed datasets using Rubin's rules.[25,28] Candidate physiological and surgical variables were excluded in turn, with those resulting in the smallest reduction in model  $R^2$  excluded first,[29] until the basic model was reached.

The  $R^2$  value from the full model was used to define a threshold for choosing a model containing only the most important additional physiological and surgical variables. The 'selected model' was the simplest model with an  $R^2$  value greater than 95% relative to the full model  $R^2$  value .

Once the selected model was finalised in the development dataset, this model was refitted using the full dataset (i.e., both developmental and validation datasets). Estimated model coefficients were reported in the form of an equation and the corresponding odds ratios (ORs) were also reported.

## Model performance and validation

Other performance measures calculated were the C-statistic, and the scaled Brier score (SBS). The C-statistic quantifies the discrimination of the model, ranging from 0.5 (non-informative) to 1 (perfect

discrimination).[30-33] The Brier score quantifies the average prediction error (accuracy) of the model predictions, compared to a naïve non-informative model. We calculated SBS ranging from 0 (non-informative) to 100% (perfect predictions).[30,34,35] Verburg et al. gave the following rule-of-thumb for interpreting a SBS for binary outcomes: <0.04 representing very weak predictions, 0.04-0.15 weak, 0.16-0.35 moderate, 0.36-0.62 strong, and >0.63 very strong.[36]

Using the development dataset, we calculated the  $R^2$  value, C-statistic, and SBS for the full model, selected model, basic model, and all models in between.

Using the validation dataset, we refitted the full model, selected model, and basic model and assessed calibration of the selected model compared to the full model and basic model by plotting observed versus predicted mortality by deciles of risk. We also calculated the  $R^2$  value, C-statistic, and SBS for the three models.

Using the full dataset, we used funnel plots to visually explore hospital trust-level variation in 90-day mortality for the basic and selected models to determine whether variation between hospital trusts was greater than expected by chance alone.[37] A hospital trust is an organisational unit in the English NHS that can include more than one hospital.

For sensitivity analysis, we produced estimates of adjusted ORs for the selected model using complete case analysis instead of using multiple imputation. A further sensitivity analysis explored using post-operative measures of actual operative severity, peritoneal soiling, and intraoperative blood loss instead of their pre-operative estimates. We used Stata 17.0 for all linkage, imputation, and analysis of data[38] and reviewed the TRIPOD guidelines for reporting on multivariable prediction models (Appendix B).[39] This work has also been reported in line with the STROCCS criteria.[40]

## Results

Overall, 846 of 10,578 (8.0%) patients died within 90 days of surgery, with 90-day mortality in the development and validation datasets of 8.2% and 7.6% respectively (Appendix Table A.3). 10,441 (98.7%) of the analysis cohort linked to the disease-specific dataset (NBOCA), the treatment-specific dataset (NELA) or both (Appendix Figure A.1). 5,803 (54.9%) patients linked to both the disease-specific and treatment-specific datasets, providing information on patient and tumour characteristics as well as physiological and surgical measures.

Table 1 shows the patient and tumour characteristics in the basic model and the candidate physiological and surgical measures, and the source of information for each variable. Where information on a variable was available from multiple datasets, reconciliation of conflicting information was undertaken (Table 1).[9]

Patient and tumour characteristics, physiological and surgical measures, and 90-day mortality are summarised in Tables 2 and 3 for the whole analysis cohort. The missing categories in the tables include patients with missing data due to non-linkage and, to a lesser extent, patients with missing values in the datasets. 55.5% of the analysis cohort had at least one missing variable. Distributions of patient characteristics were similar for the development dataset and the validation dataset (Appendix Table A.3).

The selected model included seven physiological and surgical measures (pulse rate, systolic blood pressure, breathlessness, sodium, urea, albumin and prediction of peritoneal soiling) in addition to the ten patient and tumour characteristics in the basic model (calendar year of surgery, age, sex, ASA grade, TNM T stage, TNM N stage, TNM M stage, cancer site, number of comorbidities, emergency admission).

Figure 1 shows model performance measured at each stage of the variable selection process, using the development cohort only. More details can be found in Appendix Table A.4. The full model (all patient, tumour, surgical, and physiological variables) had an  $R^2$  of 0.235 (blue markers). The percentage decrease in  $R^2$  is shown for the remaining model after each variable is removed until we are left with the basic model (patient and tumour characteristics only).

Of all physiological and surgical measures, serum albumin was the most predictive, contributing most to the  $R^2$ , therefore being the last variable removed in the variable selection process. The selected model had an  $R^2$  value of 0.225, 95.7% relative to the full model, whilst the basic model had an  $R^2$  value of 0.163, 69.2% relative to the full model (Figure 1). The selected model had a moderate SBS of 17.6%, [36] slightly lower than the full model SBS (18.6%), whereas the basic model had a substantially weaker SBS (10.9%) (Figure 1). All fitted models had good discrimination in the development cohort, with C-statistics, 0.84 for the full model, 0.83 for the selected model, and 0.80 for the basic model (Figure 1).

The measures of model performance ( $R^2$ , C-statistic, HL p-value, SBS) for the full model, the selected model, and the basic model in the validation cohort (Table 4) were slightly lower than those in the development cohort. Discrimination was lower for the basic model in the validation cohort (0.78)

whilst the full model and selected model had similar discrimination as in the development cohort (0.84 and 0.83 respectively).

Figure 2 displays calibration of the full model, selected model, and basic model using data from the validation cohort, by plotting observed mortality versus the model predicted risk by deciles of predicted risk. All three models demonstrate good calibration. There was no evidence of poor fit as HL test p-values were large for all three models (Table 4).

Figure 3 explores whether the basic model lacks calibration because it does not include the seven physiological and surgical measures included in the selected model. It displays observed and predicted mortality, using the basic model and the validation cohort, by categories of the physiological and surgical measures that were included in the selected model. The basic model had poorer calibration for patients with abnormal levels of serum albumin, urea, systolic blood pressure, or history of breathlessness at rest. On average across imputations, 8.3% of patients in the validation cohort had abnormal levels in at least one of these.

Table 5 gives adjusted ORs and 95% confidence intervals (CIs) calculated using the selected model, for all variables included in the selected model. Appendix Table A.5 gives the equation for the selected model using the whole analysis cohort. As the effect of a continuous risk factor on mortality is not easily expressed when the relationship is non-linear, Table 5 presents ORs for selected values of the continuous factors. The adjusted odds of death within 90 days of major resection increased with worse breathlessness history, lower systolic blood pressure, higher pulse rate, higher urea levels, and lower albumin. The adjusted odds of death with serum albumin levels of 25 g/L was 1.66 (CI: 1.48, 1.86) times the adjusted odds at serum albumin 35 g/L (reference value). The adjusted odds of death with urea levels of 20 mmol L<sup>-1</sup> or 30 mmol L<sup>-1</sup> were 1.57 (CI: 1.32, 1.88) and 2.18 (1.54,3.10) times the adjusted odds at urea 10 mmol L<sup>-1</sup> (reference value) respectively.

Funnel plots of adjusted 90-day mortality by hospital trust, risk-adjusted using (i) the basic model and (ii) the selected model, are shown in Figure 4. Using the selected model, five hospital trusts were above the inner limit (i.e., identified as having outlying performance). If instead the basic model was used for risk-adjustment, one of these hospital trusts would not have been identified as having outlying performance (a 'false negative') and one further hospital trust would be incorrectly identified as having outlying performance (a 'false positive').

Fewer patients would be classified as high risk and therefore receive care recommended for high-risk patients using the selected model compared to the basic model (22.9% of patients had predicted risk >0.1 and 18.5% had predicted risk between 0.05 to 0.1 using the selected model, versus 25.4% and

21.1% respectively for the basic model). The selected model classified 15.2% of patients into a lower risk category than the basic model, and 7.8% into a higher risk category.

In sensitivity analysis, a similar pattern of associations between risk factors and mortality was seen in the complete case analysis, only with greater uncertainty (Appendix Table A.6). Furthermore, results were not sensitive to using post-operative surgical measures in variable selection instead of pre-operative estimates of operative severity, peritoneal soiling, and intraoperative blood loss, and prediction did not improve substantially (results not shown).

Appendix Table A.7 compares characteristics of patients according to which datasets they were linked between. Patient characteristics were broadly similar, comparing those linked from the administrative dataset (HES) to only the disease-specific dataset (NELA), only the treatment-specific dataset (NBOCA), those linked to both, and those linked to neither. However, the patients that only linked between the spine (HES) dataset and the treatment-specific (NELA) dataset had much higher mortality compared to other groups.

## Discussion

We have developed and validated an accurate and relatively simple risk model for patients with colorectal cancer (CRC) undergoing emergency surgery, that can be used for risk-adjustment or clinical care planning. We identified seven important physiological and surgical measures which we recommend should be included along with patient and tumour characteristics to accurately predict post-operative mortality. These measures are either easy to capture (pulse rate, blood pressure, breathlessness history) or estimate (peritoneal soiling), or are completely objective (urea, albumin, sodium). This extended risk model, including these additional physiological and surgical measures, allows more accurate prediction than available basic risk models, particularly for patients with abnormal values of these additional measures.

### Comparison to other risk models

All seven of the physiological and surgical measures included in our selected model have been found to be important predictors in other models, including the surgical risk score P-POSSUM.[41,42] The discrimination of our selected model compares very favourably to existing risk models (which have c-statistics ranging from 0.732 to 0.861 [7,8,15-21] whilst including a relatively small set of risk factors that are all routinely available in patients undergoing emergency CRC surgery.

Of the seven measures we selected, four (serum albumin, urea, and sodium with breathlessness history) are composite markers of pre-existing chronic disease, and impact of current acute status. For example, breathlessness could be attributable to either cardiac or respiratory disease and cardiorespiratory insufficiency (poor functional capacity) is a known major risk factor for poor surgical outcome.[43] Breathlessness may also be a symptom of metabolic acidosis in an unwell patient with sepsis. Urea is a biomarker for kidney function, but is also affected by diet, dehydration, and proximal colonic bleeding.[44] Serum albumin falls in the acute phase response of sepsis but may also be a marker of chronic malnutrition.

### Strengths, limitations and opportunities for further work

The model was developed in a large representative national study using linked electronic health records, and candidate variables were drawn from the literature. The variables identified by the selected model are all recorded as close as possible to the time that a patient was booked for theatre. During the development of our selected model, we included the physiological measures as continuous variables. Other risk models have categorised these variables, which leads to loss of information that could mask or exaggerate relationships.[45]

We split the data into development and validation datasets by date of surgery (temporal validation), since this provides a stronger test of the validity of the predictions than splitting the data at random.[46] External validation of the selected risk model using data from other emergency CRC surgery populations would further increase confidence in predictive ability.[47]

Our study period ends in November 2019, meaning that all patients in the cohort had their emergency CRC surgery before the start of the Covid-19 pandemic. During the early pandemic period, there were rapid changes in national guidelines for cancer services. and it has been shown that post-operative mortality increased for patients having emergency CRC surgery.[48] Due to these reasons, we decided against using data from the early pandemic period for model development and validation. The selected risk model should be recalibrated using data from after the recovery of cancer services in England, once this data becomes available.

We used logistic regression to model risk of mortality. Machine learning is an alternative option, but a comparison of methods is not the focus of the current paper. Rigorous comparisons of machine learning and logistic regression have found similar performance in patients with CRC and in other patient groups. For example, a study comparing logistic regression and machine learning models to predict mortality after hospital admission using the same large national datasets demonstrated that

their performance was comparable.[49] This is fully in line with a recent systematic review that found no evidence of superior performance of machine learning over logistic regression, when both were applied according to recommended analysis strategies.[50] Therefore, it is very unlikely that a comparison between statistical and machine learning methods here would provide additional clinical insight into the risks for colorectal cancer patients undergoing emergency surgery.

We have not accounted for clustering by hospital trust in our model. In order to develop a risk model that accounts for clustering, we would need to implement this into multiple imputation, which adds considerably to the complexity of the procedure.[51] Further, since the within-hospital trust intraclass correlation coefficient was estimated as 3.8% (95% CI: 1.2, 10.8) for the selected model using complete case analysis, accounting for clustering would have little impact on the model intercept or the weights attached to risk measures.

We used spine linkage to construct the analysis cohort used here. In a recent paper comparing spine linkage with using all pairwise linkages, we have shown that this spine approach is appropriate in this setting.[9]

In this study, a substantial proportion of patients had missing data, the majority of which was due to non-linkage. We used the same multiple imputation procedure for all missing data. Although we did not treat missingness in datasets differently to missingness due to non-linkage, we did compare characteristics of patients by linkage group to assess potential for bias due to non-linkage. Patients linked only between the administrative spine dataset and the treatment-specific dataset had higher mortality. Multiple imputation methods result in unbiased estimates of model weights, provided all important variables associated with missing values are included in the imputation models.[22]. By also including mortality in these imputation models, we allowed for differences in missing measurements between survivors and non-survivors, which is much more plausible than assuming that missing measurements are similar in survivors and non-survivors. It is also important to note that a sensitivity analysis, excluding patients with missing values regardless of the reason for missingness, produced a similar pattern of results, which provides further evidence for the validity of our approach.

## Implications

This new accurate and relatively simple risk prediction model for patients undergoing emergency CRC surgery is recommended for risk-adjustment and to aid clinical care planning within the context of shared decision-making with patients and their families. The model is ready to be translated into a risk calculator that can be used to predict pre-operative risk for an individual patient. Since HES, the

administrative dataset used as the spine dataset to define our patient cohort, contains information on hospital admissions for the whole country,[11] there was no selection of patients other than the eligibility criteria described in the methods section. This should mean that our risk model is transportable to other healthcare settings with similar patients and populations. The model should be recalibrated periodically and also before being applied in new settings because overall mortality is likely to change over time and to differ by healthcare setting. For settings with very different prevalence, the weights for different factors should be reliable but the intercept may need to be adjusted to ensure good overall calibration.

Inclusion of the seven physiological and surgical measures identified in our study in the risk prediction model improves the accuracy of prediction. This will for example help to ensure that the patients at the highest risk can be identified so that they can benefit from appropriate interventions such as the presence of a consultant surgeon and anaesthetist in theatre and direct transfer to critical care in the post-operative phase.

Whereas patient and tumour characteristics are routinely available in national clinical datasets, this is not usually the case for physiological and surgical measures. Thus, for accurate risk-adjustment of CRC patients undergoing emergency surgery, it is important either to link to existing data, capture existing hospital data into databases, or to collect the key physiological and surgical measures identified in our model.

We recommend using multiple imputation to handle missing data when using the risk model for case-mix adjustment for a cohort of patients. When using the risk model for pre-operative care planning for an individual patient, it may be appropriate to assume the highest risk values of variables that are unknown.

Although the purpose of this exercise was to develop a model for risk-adjustment or clinical care planning, the findings that there are some potentially modifiable risk factors for survival may have implications for future research. However, we have only established associations and predictive ability. Further work is needed to investigate whether modifying these risk factors before surgery is feasible in the emergency setting and would improve patient outcomes.

We demonstrated an approach for developing a risk model for patients undergoing emergency resection for CRC for which there is an intersection of two sources of national clinical electronic health records, and where data linkage can be used to obtain all of the required risk factors. We included approaches which others can follow for variable selection, dealing with missing data, and assessing bias from missing data due to non-linkage. In our study, there was information available on

the patient group of interest from a disease-based dataset and a treatment-based dataset. Similar situations in which both types of datasets may be available are for example solid organ transplantation or cardiac surgery.[52,53]

### Conclusion

We showed that including seven physiological and surgical measures which are easy to measure or predict from pre-operative imaging, in addition to patient and tumour characteristics, improve the performance of models predicting risk for patients undergoing emergency CRC surgery. Inclusion of these additional measures, in our study available through linkage of electronic health records, will lead to more accurate performance monitoring of hospitals providing CRC surgery and enhance clinical care planning for individual patients.

## Acknowledgements

We give thanks for our patient representatives for their help in defining the research questions and commenting on research outputs.

The National Bowel Cancer Audit is commissioned by the Healthcare Quality Improvement Partnership (HQIP) as part of the National Clinical Audit and Patient Outcomes Programme, and funded by NHS England and the Welsh Government ([www.hqip.org.uk/national-programmes](http://www.hqip.org.uk/national-programmes)). Neither HQIP nor the funders had any involvement in the study design; in the collection, analysis, and interpretation of data; in the writing of the report; or in the decision to submit the article for publication.

The National Emergency Laparotomy Audit is commissioned by the Healthcare Quality Improvement Partnership (HQIP) as part of the National Clinical Audit and Patient Outcomes Programme (NCAPOP). The programme is funded by NHS England, the Welsh Government and, with some individual projects, other devolved administrations and crown dependencies: [hqip.org.uk/national-programmes](http://hqip.org.uk/national-programmes).

This work uses data provided by patients and collected by the NHS as part of their care and support.

As the National Bowel Cancer Audit involves analysis of data for service evaluation, it is exempt from UK National Research Ethics Committee approval. Section 251 approval was obtained from the Ethics and Confidentiality Committee for the collection of personal health data without the consent of patients. The study was performed in accordance with the Declaration of Helsinki.

The data used in this study are available from NHS Digital and Public Health England's Office for Data Release but restrictions apply to the availability of these data, which were used under license for the current study, and so are not publicly available. We do not have permission to share the patient-level records used in our analysis.

This research was registered at ResearchRegistry.com under research registration number:

<redacted>.

## Figure captions

Figure 1: Pooled C-statistic and relative pooled R-squared for the full model, basic model, and all intermediate models. \* Scaled pooled Brier score (SBS) also given for the full model, selected model, and basic model. Using development dataset (December 2016 to November 2018).

\*The full model includes all patient and tumour characteristics, and physiological and surgical measures. The basic model includes all patient and tumour characteristics. Each intermediate model is defined by removing one of the physiological and surgical measures from the previous model, according to which would result in the smallest difference in pooled R-squared. E.g., model 2 includes all variables included in the full model except operative severity. The selected model includes all patient and tumour characteristics, and the physiological and surgical measures highlighted in bold.

Figure 2: Calibration plot showing proportions of patients that died within 90 days (observed mortality) versus mortality predicted using the full model, selected model, and basic model, by deciles of predicted risk. Using validation dataset (December 2018 to November 2019).

Figure 3: Calibration plot showing proportions of patients that died within 90 days (observed mortality) versus mortality predicted using the basic model, by categories of variables that are included in the selected model but not the basic model. Using validation dataset (December 2018 to November 2019).

Figure 4: Funnel plots of adjusted 90-day mortality risk-adjusted using the basic model and the selected model, by hospital trust. Using full dataset (December 2016 to November 2019).

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