Supporting the ambulance service to safely convey fewer patients to hospital by developing a risk prediction tool: Risk of Adverse Outcomes after a Suspected Seizure (RADOSS) – protocol for the mixed-methods observational RADOSS project

Adam J Noble, Suzanne M Mason, Laura J Bonnett, Markus Reuber, Jasmine Wright, Richard Pilbery, Richard M Jacques, Rebecca M Simpson, Richard Campbell, Alison Fuller, Anthony Guy Marson, Jon Mark Dickson

ABSTRACT

Introduction Ambulances services are as asked to further reduce avoidable conveyances to emergency departments (EDs). Risk of Adverse Outcomes after a Suspected Seizure seeks to support this by: (1) clarifying the risks of conveyance and non-conveyance, and (2) developing a risk prediction tool for clinicians to use ‘on scene’ to estimate the benefits an individual would receive if conveyed to ED and risks if not.

Methods and analysis Mixed-methods, multi-work package (WP) project. For WP1 and WP2 we shall use an existing linked data set that tracks urgent and emergency care (UEC) use of persons served by one English regional ambulance service. Risk tools are specific to clinical scenarios. We shall use suspected seizures in adults as an exemplar.

WP1: Form a cohort of patients cared for a seizure by the service during 2019/2020. It, and nested Knowledge Exchange workshops with clinicians and service users, will allow us to: determine the proportions following conveyance and non-conveyance that die or re-contact UEC system within 3 (30) days; quantify the proportion of conveyed incidents resulting in ‘avoidable’ ED attendances (AA); optimise risk tool development; and develop statistical models that, using information available ‘on scene’, predict the risk of death/re-contact with the UEC system within 3 (30) days and the likelihood of an attendance at ED resulting in an AA.

WP2: Form a cohort of patients cared for a seizure during 2021/2022 to ‘temporarily’ validate the WP1 predictive models.

WP3: Complete the ‘next steps’ workshops with stakeholders. Using nominal group techniques, finalise plans to develop the risk tool for clinical use and its evaluation.

STRENGTHS AND LIMITATIONS OF THIS STUDY

⇒ Risk of Adverse Outcomes after a Suspected Seizure will use a ‘cutting-edge’ linked data set that captures service use in one ambulance region using data high in quality and coverage.

⇒ The parameters of the outcome measures used to describe risks and the variables tested for their ability to predict these outcomes will be informed by stakeholders and service users.

⇒ The large, pseudoanonymised nature of the linked data set will require the use of a generic definition of an ‘avoidable attendance’ whose validity for suspected seizures is not yet known.

⇒ As there are no equivalent linked data sets available for other ambulance regions, the validity of the derived prediction models will need to be determined within a cohort of patients treated within the same region, but at a later date.

INTRODUCTION

Context and drive for health service innovations

Ensuring people ‘get the right care at the right time in the optimal care setting’1 is a key ambition of the UK’s National Health Service (NHS). Ambulance services have a role to play. They
should only be conveying patients to emergency departments (EDs) if it is clinically appropriate or there is no alternative service to provide safe and ongoing care.

Traditionally, UK ambulance services’ primary roles were to provide emergency call handling and transportation to hospital. However, as the nature of the calls it receives has shifted towards non-life-threatening conditions, services and the clinicians working within them have needed to evolve.

NHS England and Improvement has identified that ambulance clinicians require more support with their changing role. Certain presentations continue to be ‘over-conveyed’ and reductions in ambulance conveyance rates are stalling. At the same time, ambulance services are under pressure to provide a timely response to an increasing number of calls, while facing increased handover delays at EDs.

Strategies are thus needed to support appropriate and safe decision-making on scene that minimises avoidable conveyance. The Risk of Adverse Outcomes after a Suspected Seizure (RADOSS) project seeks to generate ways of providing ambulance clinicians with this support.

### Why is reducing clinically unnecessary conveyance important?

Clinically unnecessary conveyances to EDs result in ‘avoidable attendances’ (AAs). An AA is where the patient does not require the facilities of a type 1 ED to manage their healthcare problem. AAs can harm the patient and have implications for others since they restrict ED capacity.

Approximately 15% of ED attendances currently meet the RADOSS project’s definition of an AA. In 2021/2022, this equated to ~2.3 million visits.

Patients and the public are broadly supportive of non-conveyance. Research has identified that they are frustrated by inappropriate conveyance to ED and say assessment by an ambulance clinician itself has a therapeutic value.

Importantly, UK data indicate non-conveyance following assessment by ambulance clinicians is safe. Overall, 85% of people experience no subsequent health event within 3 days of non-conveyance (9% recontact the ambulance service, 12.6% attend ED, 6.3% are admitted and 0.3% die).

### What is known about how ambulance clinicians decide who to convey?

Systematic reviews highlight the complex nature of conveyance decisions. Factors beyond patient need can affect them. Oosterwold et al’s framework (online supplemental file 1) summarises macro, meso and micro factors. Work has started to address some of these.

However, given reductions in conveyance have stalled, other factors in the model need addressing.

### What could help clinicians identify cases suitable for non-conveyance?

Ambulance clinicians are critical of current support, saying non-conveyance guidelines and protocols are difficult to apply to the nuances of cases.

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**Table 1** The RADOSS project’s aims and objectives and the work packages that address them

<table>
<thead>
<tr>
<th>Aims</th>
<th>Objectives</th>
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<tr>
<td>(1) Calculate the risks and benefits of conveyance to hospital after a suspected seizure.</td>
<td>a. Describe the characteristics of those conveyed and those not conveyed to ED by one representative English ambulance service (WP1a).</td>
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<td>b. Compare the proportions following conveyance and non-conveyance that die and/or recontact the UEC within 3 (and 30) days (WP1a).</td>
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<td>c. Quantify the proportion of incidents conveyed to ED that meet the definition of an AA (WP1a).</td>
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<td>(2) Create a risk prediction tool that predicts the likelihood that an individual will die and/or recontact the UEC system within 3 (and 30) days if not conveyed and the likelihood that their conveyance to ED would result in an AA.</td>
<td>d. Optimise the prediction tool development by completing KE workshops with service users and ambulance and ED clinicians to get views on predictors considered for inclusion in the models, the way the outcome measures of death, UEC recontact and AA are defined and risk score presentation (WP1b).</td>
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<td></td>
<td>e. Develop statistical models to predict a person’s risk of death/recontact with the UEC system within 3 (and 30) days and the likelihood of their attendance at ED being classed an AA if conveyed (WP1a).</td>
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<td>f. ‘Temporally’ validate the predictive models using data from the same ambulance service for a later time period (WP2).</td>
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<td>(3) Establish a pathway to clinical implementation of the risk prediction tool and maximise usage of RADOSS findings.</td>
<td>g. Complete ‘next steps’ workshops with stakeholders to finalise plans to refine the tool for clinical use and its evaluation (WP3).</td>
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<td>h. Complete a proactive dissemination and knowledge mobilisation strategy (WP3, WP4).</td>
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AA, avoidable attendance; ED, emergency department; KE, Knowledge Exchange; RADOSS, Risk of Adverse Outcomes after a Suspected Seizure; UEC, urgent and emergency care; WP, work package.
The National Ambulance guidelines currently recommend that different presentations have also not been fully determined.10 38 40–41

Given this, promising ways of supporting clinicians include: (1) securing and disseminating clear evidence on the risks of conveyance and non-conveyance by presentation, and (2) providing a risk prediction tool that would allow clinicians to predict the likelihood that conveyance to ED of the individual they are caring for would result in an AA and the likelihood of them experiencing adverse health events if not conveyed. This direction aligns with recommendations by Lord Carter12 and others.4 26

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<th>Reason</th>
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<tbody>
<tr>
<td>1</td>
<td>Frequently seen</td>
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<tr>
<td>2</td>
<td>‘Over-conveyed’</td>
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<td>3</td>
<td>Redeemable cause of avoidable attendance</td>
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<td>4</td>
<td>‘Alternative care pathways’ available</td>
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<td>5</td>
<td>User preference</td>
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<td>6</td>
<td>Cost</td>
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<td>1</td>
<td>Responsible for ~211 000 ‘999’ calls per year in England; 7th most common presentation.22 43</td>
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<td>2</td>
<td>‘Almost all (~97%) receive a face-to-face ambulance response.’103</td>
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<td>3</td>
<td>Around 70% of suspected seizure cases are conveyed to ED.63 103–106 This is despite national guidelines stating most will not require ED.47</td>
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<td>4</td>
<td>Suspected seizures are dramatic and frightening and traditional training emphasises status epilepticus—a rare and life-threatening condition. However, most ‘999’ suspected seizures are low risk and persons return to their normal self without intervention.35</td>
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<td>5</td>
<td>Most of those presenting have established, treated epilepsy and have experienced an uncomplicated seizure for which they require rest and reassurance.107</td>
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<td>6</td>
<td>Seizures currently have the third highest conveyance rate of all presentations.10</td>
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<td>3</td>
<td>Clinicians identify suspected seizures as a readily redeemable cause of AAs.32 33 35 46 108</td>
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<td>4</td>
<td>Alternative care pathways are becoming available for clinicians to use.109 110 They could, unlike visits to ED,111 prompt improvements in ambulatory care and so address health inequalities.112</td>
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<tr>
<td>5</td>
<td>People with established epilepsy and those with other seizure presentations, such as non-epileptic attack disorder, usually want to avoid ED after an uncomplicated seizure;116–118 preferring to recover at home.</td>
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<tr>
<td>6</td>
<td>Unnecessary conveyance to ED puts them at risk of iatrogenic harm and overinvestigation.118–120</td>
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<td>2</td>
<td>In England, the annual cost to the NHS of unplanned hospital care for suspected seizures is ~£90 million.121</td>
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Evidence suggesting a tool predicting benefit/risk of non-conveyance could be developed

The information used by any risk prediction tool should reflect what is available to the clinician at the time conveying decisions are made (and is accessible for derivation). Ambulance clinicians do not typically have access to a patient’s full medical record. What is available is the information they record using structured fields on a patient care record (PCR) about the patient’s demographics, medical history, clinical features, physiological observations as well as details relating to the care provided. Also available is structured dispatch information. Online supplemental file 2 indicates the range of data available.

So far, only a selection of this information has been examined for its relationship to the outcomes of interest. While exploratory in nature, studies have identified that recontact with the urgent and emergency care (UEC) system and death following non-conveyance, and AAs following conveyance, are not random but more common in certain subgroups (eg, patient age, sex, time of call, day of week, presence of comorbidities and social deprivation.5 9 22 58–60). A testament to the utility of the information available to ambulance clinicians is Patton and Thakore’s61 study findings. ED clinicians reviewed ambulance PCRs of patients conveyed to ED and identified those whose attendances they suspected would be AA. This was then repeated when ED clinicians had access to the PCR data

What is a risk prediction tool?

Risk prediction tools use ≥2 pieces of patient data to generate a personalised estimate of the likelihood that an individual will experience a certain endpoint within a specified time frame. Currently, there are no prediction tools relating to non-conveyance.26 However, evidence suggests they could be developed (see Evidence suggesting a tool predicting benefit/risk of non-conveyance could be developed).

Ambulance clinicians already use such tools to predict other outcomes (eg, ref 43–45) and they want more.40 46 The National Ambulance guidelines currently recommend 11 such tools (none relate to seizures). Risk prediction tools do not replace clinical judgement but support it. There is evidence they can improve patient outcomes and satisfaction and avoid unnecessary care.48–53

Methodological standards exist48 for their development. To facilitate uptake and sustained use, their development needs to be carefully informed by the views of intended users.55 56 There is no single pathway by which a tool enters practice, but good practice states confirmation be obtained that it provides valid predictions on a sample different (in time or place) from the one used for model derivation.57
and ED notes. Clinicians were confident in identifying AAs on the basis of the PCR alone.

**Current project**

**Overview and aims**

To address the identified needs and information gaps, the 24-month mixed-methods RADOSS project is being completed. It has the following aims: (1) calculate the risks and benefits of conveyance to hospital after a suspected seizure; (2) create a risk prediction tool that predicts the likelihood that an individual will die and/or recontact the UEC system within 3 (and 30) days if not conveyed and the likelihood that their conveyance to ED would result in an AA; and (3) establish a pathway to clinical implementation of the risk prediction tool and maximise usage of RADOSS findings. The project’s related objectives were noted in table 1.

Risk prediction tools are specific to clinical scenarios. We are therefore focusing on patients experiencing suspected seizures. Seizures are a topic of interest in their own right but also an ideal exemplar since they are frequently encountered by the service and ‘over-conveyed’. Table 2 expands on the reasons.

RADOSS consists of four work packages (WP). WP1 is the main one. It involves a cohort study (WP1a) and a Knowledge Exchange (KE) study (WP1b). WP2 is smaller and focuses on validation via a second cohort study. WP3 focuses on ‘next steps’ on the journey to implementation of the tool in the NHS, and WP4 on dissemination (figure 1). According to Greene et al’s conceptual framework, the purpose of using a mixed-methods approach is both ‘development’ and ‘expansion’.

**Routine data source: cured+**

For WP1a and WP2, we will use a cutting-edge database called ‘CUREd+’. Currently being developed by the Centre for Urgent and Emergency Care Research, it will map UEC use by individuals served by the Yorkshire Ambulance Service (YAS) from 2011 to 2022. It contains records of all ambulance contacts and these are linkable to any subsequent ambulance, hospital (ED, inpatient) and death records (Office for National Statistics (ONS) mortality register). Further information is provided in table 3.

**METHODS AND ANALYSIS**

**Work package 1**

WP1a: retrospective cohort study 1

**Purpose**

- Describe the pattern of calls for suspected seizure, the type of ambulance responses received and the characteristics of the patients accounting for them.
- Determine and compare the rate of death and recontact with the UEC system of those seen by the ambulance service for a suspected seizure who were and were not conveyed to ED.
- Determine the proportion of suspected seizure incidents conveyed to ED that resulted in an AA: develop predictive models for risk of death/recontact with the UEC system within 3 (and 30) days following conveyance and non-conveyance and risk of attendance at ED being classed an AA if conveyed.

**Figure 1** Summary of RADOSS project. Using four WPs, we will develop a risk prediction tool for people after a suspected seizure; we will validate the tool, plan its implementation and disseminate the findings. ED, emergency department; NHS, National Health Service; RADOSS, Risk of Adverse Outcomes after a Suspected Seizure; WP, work package.
Combine the predictive models to form a draft tool that can potentially provide estimates of an individual’s risk of death/recontact with the UEC system if managed by non-conveyance; risk of death/recontact with the UEC system if managed by conveyance; and the risk of their attendance at ED being classed as an AA if conveyed.

To do this, a retrospective cohort of adults cared for a suspected seizure by YAS will be studied.

Identification

Index events will be identified by searching CUREd+ for persons managed by YAS for a suspected seizure between 1 February 2019 and 31 January 2020. Eligibility criteria are presented in table 4.

The unit of analysis will be the patient, with the first recorded episode being the index event and subsequent episodes ≤3 days defined as recontacts (or 30 days for the secondary analysis).

Data extract

The data extract provided will include any ambulance, ED (Emergency Care Data Set; Hospital Episode Statistics (HES) Accident and Emergency (due to overlap in system use)), urgent inpatient (HES Admitted Patient) and death (ONS) records that relate to the index events which started within 30 days.

Outcome measures

Death/recontact with the UEC system following ambulance care and the likelihood of an AA occurring if conveyed to ED are important outcomes to clinicians and service users. Below we describe how the index events will be classified according to these two measures.

Measure 1 (safe/unsafe: death or recontact with UEC)

All index events, both conveyed and non-conveyed to ED, will be classified according to whether linked data indicate the patient involved died and/or recontacted the UEC (defined as any ambulance, ED or unscheduled inpatient care).

For the primary analysis, we propose a time frame of up to 3 days from the event within which death must have occurred or recontact started. This has been specified by paramedics and other stakeholders. It aligns with evidence that when considering all ambulance presentations, ~75% of deaths/UEC recontacts following non-conveyance occur within 3 days. We shall though still confirm its suitability with clinicians and service users via WP1b. For secondary analyses, a time frame of 30 days is proposed.

Deaths within the cohort should be rare. Nonetheless, when describing and using deaths we shall report them with and without exclusion of persons where death was associated with end-of-life care.

Measure 2 (avoidable/unavoidable ED attendance)

Index events that resulted in conveyance to ED will be classified according to whether they resulted in an AA or not.

To determine this, the events will be assessed against O’Keeffe et al’s9 definition. Namely, a person has been involved in an AA if routine hospital coding for the attendance indicates it did not result in the person being investigated (except urinalysis, pregnancy test, dental investigation) or treated (except prescription, recording vital signs, dental treatment or guidance/advice), and they were discharged.

O’Keeffe’s system has advantages. It is generic, applicable to all ages, based on process of care rather than initial triage score and has been adopted by the NHS. It is also quick and routine data have been found to be sufficient to mean it can be applied to ~98% of attendances. A possible disadvantage is it assumes all investigations, treatments and admissions were clinically indicated.
Table 4  Participant inclusion and exclusion criteria for different WPs

<table>
<thead>
<tr>
<th>WP</th>
<th>Inclusion criteria</th>
<th>Exclusion criteria</th>
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| WP1a: retrospective cohort study 1 | ▶ Incident cared for by YAS.  
▶ Person aged ≥16 years (no upper age limit).  
▶ Incident occurred between 1 February 2019 and 31 January 2020.*  
▶ ‘Chief complaint’ (or other complaint) selected by attending ambulance clinician on PCR was ‘convulsions/fitting/seizure’ OR, if empty, call handler coded it as AMPDS protocol 12 (‘Convulsions’).  
▶ No restriction on type of ambulance response incident received (ie, could have been ‘Hear & Treat’, ‘See & Treat’, ‘See & Convey to ED’ or ‘See & Convey elsewhere’†). | ▶ The <1% of events coded as AMPDS02. These relate to seizures in someone potentially pregnant. (Guidelines state these should be conveyed because of eclampsia risk.†)  
| WP1b: Knowledge Exchange (KE) workshops | ▶ Aged ≥18 years (no upper age limit).  
▶ Attended to by an ambulance during prior 12 months for a suspected seizure/s OR a significant other to such a person (eg, family member, friend).  
▶ Incident could be related to epilepsy, non-epileptic attack disorder or syncope. (They account for ~70% of events.†)  
▶ Able to provide informed consent and participate in a workshop independently in English. | ▶ Severe current psychiatric disorders (eg, acute psychosis).  
▶ Terminal medical condition.  
▶ Severe current psychiatric disorders (eg, acute psychosis).  
▶ Terminal medical condition. |
| Clinicians          | ▶ Aged ≥18 years (no upper limit).  
▶ Ambulance clinician, ED doctor or nurse.  
▶ Works in England.  
▶ Able to provide informed consent and participate in a workshop independently in English. | |
| WP2: retrospective cohort study 2 | ▶ Incident cared for by YAS.  
▶ Person aged ≥16 years (no upper age limit).  
▶ Incident occurred between 1 July 2021 and 30 June 2022.†  
▶ ‘Chief complaint’ (or other complaint) selected by attending ambulance clinician on PCR was ‘convulsions/fitting/seizure’ OR, if field was empty, call handler coded it as AMPDS 12.  
▶ No restriction on type of ambulance response incident received. | ▶ The <1% of events coded as AMPDS02. These relate to seizures in someone potentially pregnant. |
| WP3: ‘Next Steps’ workshops | ▶ Aged ≥18 years (no upper limit).  
▶ Ambulance clinician, seizure/epilepsy guideline developer, user group representative, seizure specialist (eg, neurologist/epilepsy nurse), commissioning representative.  
▶ Able to provide informed consent and participate in a workshop independently in English.  
▶ Lives in UK. | ▶ Severe current psychiatric disorders (eg, acute psychosis).  
▶ Terminal medical condition. |

*The time period does not include periods of industrial action; is before changes in use of acute services due to COVID-19 became apparent128; and is before the first UK COVID-19 fatality.129  
†These are the labels used by the National Health Service (NHS) to record the main types of responses that ambulance services provide to incidents. Further detail is available from NHS England.130  
‡This time period represents the most contemporary 12 months for which linked data will be available. It excludes COVID-19 national ‘lockdowns’ and the start aligns with when most COVID-19 legal restrictions in England were removed.  
AMPDS, Advanced Medical Priority Dispatch System; ED, emergency department; PCR, patient care record; WP, work package; YAS, Yorkshire Ambulance Service.

Some may have happened for other reasons (eg, routine or inappropriate administration of test). Thus, we shall describe the reasons why any WP1a cases satisfied the criteria for an unavoidable attendance. Moreover, via WP1b, we shall ask ED clinicians to what extent suspected seizure cases attending their EDs could satisfy the criteria of an unavoidable attendance based on routine practice. Should it prove warranted, a sensitivity analysis will be conducted with and without such cases.

Sample size

Predictive models for the (1) risk of death/UEC recon- 
tact following conveyance, (2) risk of death/UEC recon- 
tact following non-conveyance, and (3) risk of an AA following conveyance could be developed. To permit robust testing of at least 40 candidate predictor parameters for each of these models, Riley et al.74 formulae using standard parameters indicate: for model (1), a need for 2567 index events, with 103 experiencing the target event; for model (2), a need for 2194 index events, with 308 experiencing the target event; and for model (3), up to 2194 index events, with 461 experiencing the target event. Twelve months of YAS data should be sufficient to satisfy these requirements. Online supplemental file 3 details the reasons why and provides further information on the sample size calculation.

Data management and analysis

Curation

A statistician, with support from a data manager, will complete data quality checks on the data extract, identifying missing and incongruent values.

Describing sample and patient outcomes

The characteristics of the calls for suspected seizures (dispatch codes, time of day, day of week, location), the patients accounting for them and the ambulance response they receive (proportions managed by ‘Hear &
Table 5  Topic guide areas that WP1b Knowledge Exchange workshops will explore (emphasis will vary depending on whom the workshop is for)

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| 1    | Potential predictors | ► Asked for views on potential predictors, including perceived utility, reliability, validity and consistency in measurement. 
| 2    | Parameters of outcome measures | ► Asked whether any routine ED practices could mean seizure cases by default would not satisfy AA definition.  
► Asked about any known differences between EDs and hospitals in how codes for incidents are applied at them that could undermine validity of definition that is based on them. 
► What time frame for death and recontact with UEC would be most supportive for conveyance decisions. 
| 3    | Optimal way to present risk scores | ► Asked whether percentage probability and/or broad risk categories wanted, whether visual aids would help and if estimates of uncertainty around probabilities wanted.  
► Illustrations of options offered. 
► Asked what ‘low’, ‘medium’ or ‘high’ risk of death, UEC recontact or AA would look like to them in percentage terms. 
| 4    | Optimal format for tool | ► Asked how they might want such a tool to be presented in future (eg, web tool, nomogram, graphical score chart), who should have access to it and the extent to which they would want it integrated into existing workflows. 

AA, avoidable attendance; ED, emergency department; UEC, urgent and emergency care; WP, work package.

Treat’, ‘See & Treat’, ‘See & Convey to ED’ and ‘See & Convey elsewhere’) will be described. For events receiving the response ‘See & Convey to ED’, we shall:
► Tabulate ED discharge diagnoses.  
► Calculate the proportion satisfying the AA definition.  
► Tabulate the reason/s why persons did not satisfy the AA definition.  
► Calculate the proportion recontacting the UEC system within 3 (and 30) days (with and without inclusion of those whose subsequent contact/s meet the AA definition).  
► Calculate the proportion dying within 3 (and 30) days and reasons.  
For events receiving a face-to-face response but not conveyed to ED (ie, ‘See & Treat’, ‘See & Convey elsewhere’), we shall:
► Calculate the proportion recontacting UEC system within 3 (and 30) days (with and without those whose subsequent contact/s meet the AA definition; also, with and without those originally non-conveyed to ED because they refused).  
► Calculate the proportion dying within 3 (and 30) days and reasons.  

Derivation of prediction models and management of missing data
As the outcome measures are binary, multivariable logistic regression will be used to derive the predictive models. Reporting will be done according to best practice. The pool of candidate predictors for testing will be informed by WP1b (see WP1b: KE workshops) and chosen based on clinical relevance, consistency in measurement and ease of use in practice. Where possible, variables will be used in their original form.

While missingness on core data items is anticipated to be low, missingness on wider items might be higher since tests may not be performed if expected to be normal and not all PCR fields are mandatory. Where data are ‘expectedly’ missing (ie, the test is not performed as not clinically indicated), an additional category of ‘not clinically indicated’ will be added to the variable. In the case of more than 10% missingness for any other variable, multiple imputation via chained equations will be undertaken. A set of 20 imputed data sets will be created using predictive mean matching. Functional form for continuous variables will be assessed via fractional polynomials of more than 10% missingness for any other variable, multiple imputation via chained equations will be undertaken. A set of 20 imputed data sets will be created using predictive mean matching. Functional form for continuous variables will be assessed via fractional polynomials within each imputed data set. Variables will be selected for inclusion in the final model within each imputed data set via backward selection with a p value of 0.10. Variables that feature in at least 10 of the 20 imputed models will be selected for the final model. Pooled OR and intercepts will be calculated according to Rubin’s rule.  

Apparent measures of model performance will be calculated for the final multiple imputed model. The area under the receiver operating characteristic curve will be calculated to assess the final model’s discriminative performance. Discrimination refers to the ability of the prognostic model to differentiate between those who experienced the event and those who did not. We will report the calibration slope and the ratio of expected to observed events to evaluate calibration, how closely the probability of the event predicted by the model agrees with the observed probability. C-statistics resulting from the imputed data set will be pooled via robust methods and therefore the median of the imputed estimates will be presented. Calibration will also be observed via a calibration plot for each imputed data set separately and the median of the imputed estimates provided. To account for sampling variability and enable adjustment of the regression coefficients for overfitting, the final model will be internally validated via bootstrap resampling. In each of 500 bootstrap samples, the entire modelling process, including predictor selection, will be repeated and the apparent model performance (calibration and discrimination in the bootstrap sample) will be compared with the performance in the original sample per multiple imputed data set. The median optimism...
across all imputed samples will then be used to calculate the optimism-adjusted C-statistic and optimism-adjusted calibration slope.\textsuperscript{84} Using the latter as a uniform shrinkage factor, all the predictor effects in the final developed model will be penalised in order to account for overfitting.\textsuperscript{85}

The pool of potential predictors for the backward selection will be any predictor in a final multivariable model for each imputed data set.

**Combining the predictive models to form a draft tool**

The three derived models will be combined to form a single, Excel-based draft version of a tool that seeks to provide estimates of an individual’s risk of death/recontact with the UEC system if managed by non-conveyance; risk of death/recontact with the UEC system if managed by conveyance; and the risk of their attendance at ED being classed as an AA if conveyed. The manner in which it is presented will be informed by WP1b and previous work by Bonnett et al.\textsuperscript{86} Examples of tools that have combined predictive models to provide clinicians with different estimates to inform decisions include the CHA2DS2-VASc/HAS-BLED\textsuperscript{87} and the cancer PREDICT tool.\textsuperscript{88}

**WP1b: Ke workshops**

**Purpose**

- Optimise prediction tool development by completing KE workshops with service users, ambulance clinicians and ED clinicians to get views on candidate predictors, the way the outcome measures of death, UEC recontact and AA are defined and risk score presentation.

**Design**

KE workshops will be run online using videoconferencing technology. Wilkins and Cooper\textsuperscript{89} defined KE as a two-way exchange between researchers and research ‘users’ to share ideas, evidence, experiences and skills. It goes beyond telling people things and is a process of listening and interaction, with a goal to generate mutual benefit.

**Participants**

**Service users**

Purposive sample of ~20–30 persons recently receiving ambulance care for a suspected seizure/s and their significant others. Full eligibility is presented in table 4.

Individuals shall be recruited via user groups affiliated with the different conditions (including epilepsy deaths). They shall circulate advertisements directly to their members and within publications.

**UEC clinicians**

Sampling will be purposive, consisting of a group of ~20–30 informed individuals/‘experts’ deemed to have high professional knowledge and clinical experience of the UEC system.

The national ‘Lead Paramedic Group’ will circulate advertisements, with priority being given to ambulance clinicians from the n=6 services that have used Advanced Medical Priority Dispatch System. To recruit ED clinicians, the Royal College of Emergency Medicine Yorkshire and Humber regional board shall circulate advertisements.

**Procedure**

Workshops for service users and ambulance clinicians will run separately. To maximise participation, we anticipate two to three for each. They will be conducted by a qualitative researcher. For those with clinicians, statistician LJB will assist.

Workshops will start with an explanation of the risk tool, aims and a presentation of the potential predictors and proposed outcome measures. A topic guide will direct the conversation. It will be finalised on the basis of the literature,\textsuperscript{86,88} our experience and key uncertainties regarding the tool’s future implementation surfaced by completion of Greenhalgh et al’s Non-adoption, Abandonment, Scale-up, Spread and Sustainability Complexity Assessment Tool Long.\textsuperscript{91} The main areas that the workshops intend to cover are shown in table 5. Workshops will last ~60–90 min.

**Analysis**

Data will include field notes and audio recordings. A qualitative researcher, supported by the wider team, will take an inductive and deductive approach to analysis. NVivo will provide a transparent account of the work. Nodes (codes) will be created to mark relevant concepts and topics in the documents. Lower level nodes will be grouped into themes.

**Work package 2**

**WP2: Retrospective cohort study 2**

**Purpose**

- ‘Temporally’ validate WP1a’s predictive models.

The predictions of the WP1a models will be tested on a data set relating to patients cared for by YAS during a 12-month time period different from that used for derivation.

**Identification, data linkage, data checks and outcome measures**

CUREd+ will be searched to identify events as done for WP1a, except the date range will be 1 July 2021 to 30 June 2022 (table 4). Outcome measures and processes used will be the same.

**Sample size**

The validation sample will be similar to that used for derivation. It will thus satisfy the recommendation that validation samples include ≥200 cases experiencing the target events.\textsuperscript{82,83}

**Data management and analysis**

**Describing sample and patient outcomes**

Sample contributing data will be described as for WP1a.

**Comparison with time period used for model derivation**

Number of calls for and the characteristics of the patients presenting with suspected seizures during the derivation and validation periods will be compared, as will the proportions...
Table 6 Dissemination actions (in addition to WP3)

<table>
<thead>
<tr>
<th>Activity</th>
<th>Detail</th>
</tr>
</thead>
<tbody>
<tr>
<td>1 Promoting awareness/engagement</td>
<td>Notification of the project’s funding and progress sent to medical directors and lead consultant paramedics of all ambulance services, National Clinical Director for Urgent Care for NHS England, National Ambulance Commissioners Network; National Ambulance Urgent and Emergency Care Group subgroup of the Association of Ambulance Chief Executives, National Ambulance Research Steering Group.</td>
</tr>
<tr>
<td>2 Interim updates</td>
<td>As project progresses, accessible briefings are produced and disseminated to funders; stakeholders; service user groups; policy makers; NHS audiences; and research bodies. Include NHS Improvement and NICE who identified need for such research.</td>
</tr>
<tr>
<td>3 Peer-reviewed outputs</td>
<td>Minimum of 2 papers in peer-reviewed journals which would appeal to clinical, organisational, general health and social policy audiences.</td>
</tr>
<tr>
<td>4 Taking evidence to practitioners</td>
<td>Findings circulated via NHS network newsletters, in practitioner journals and general press.</td>
</tr>
<tr>
<td>5 Taking evidence to clinicians</td>
<td>Oral and poster presentations at neurology and acute/emergency care conferences and fora.</td>
</tr>
<tr>
<td>6 Taking evidence to participants</td>
<td>Summary of project’s findings distributed to participants in the different WPs.</td>
</tr>
<tr>
<td>7 Media briefings</td>
<td>Updates on websites including YAS, Epilepsy Action and universities.</td>
</tr>
<tr>
<td>8 Taking evidence to service users</td>
<td>Service users and significant others/carers will clearly be interested in study outcomes. Epilepsy Action will feature study with patient experience stories in communications with epilepsy community.</td>
</tr>
</tbody>
</table>

NHS, National Health Service; NICE, National Institute for Health and Care Excellence; WP, work package; YAS, Yorkshire Ambulance Service.

conveyed to ED, the proportions whose attendance meets the AA definition and the proportions dying/recontacting the UEC within 3 (and 30) days. Differences will be described and tested for statistical significance.

Temporal validation of predictive models
Predictors and regression coefficients from the final internally validated, optimism-adjusted models will be applied to the WP2 data set to predict the target outcomes. The performance of the models will be quantified by comparing predictions with observed outcomes. Performance will be assessed using measures of discrimination and calibration. Model recalibration will be undertaken if there is systematic underprediction or overprediction.

Work package 3
Next steps’ workshops

Purpose
- Finalise plans to refine the risk tool for clinical use and its evaluation.
- If the developed models are found to make predictions with an acceptable level of validity then we would have satisfied the requirements for the tools use within practice. We would therefore need to finalise its presentations for clinical use and evaluate its impact on clinical practice. To ensure any plans for this are acceptable to stakeholders and address their information needs, ‘Next steps’ workshops will be completed.

Design
Two online workshops, each lasting ~3 hours. We shall limit each to approximately eight to nine participants.

Workshops will start with a presentation of RADOSS findings and our draft ‘next steps’. To secure stakeholders’ views of these we would use an adapted version of the nominal group technique.

With respect to what evaluation we propose we consider it appropriate to make this judgement nearer the time. A cluster randomised controlled trial would likely be most rigorous. However, various factors can influence and constrain design choice. This includes time frame within which evidence is required and regulations at the time surrounding risk tools.

Recruitment
We shall seek representation from:
- Care guideline providers (via Joint Royal Colleges Ambulance Liaison Committee panel for seizures; National Institute for Health and Care Excellence panel for epilepsy).
- User groups (including Epilepsy Action, Epilepsy Society, FND Action, SUDEP Action and others).
- Seizure specialists (via International League Against Epilepsy; Epilepsy Specialists Nurses Association).
- Commissioners (via National Ambulance Commissioners Network).

Personal invitations will be sent. To maximise attendance, we shall exploit existing relationships our team has. We shall overinvite by ~30%. Table 4 provides the eligibility criteria.

Procedure
Workshops will be facilitated by the investigative team. Presentations will be pre-recorded to reduce opportunity for technical difficulties.
**Analysis**

Field notes will be kept. Delegates’ involvement will be anonymous. A summary of the findings will be generated and discussed by the investigators and the ‘next steps’ plan finalised.

**Work package 4**

**Purpose**

- Disseminate findings to key stakeholders and maximise evidence usage.

**Dissemination and outputs**

We shall engage in a proactive dissemination and knowledge mobilisation strategy to ensure those who are considering developing, funding or supporting non-conveyance strategies are aware of the project and its findings. All investigators shall contribute, and the media departments of involved institutions shall help. As well as conducting WP3, dissemination will consist of the items in table 6.

**DISCUSSION**

**Patient and public involvement**

This research was instigated by evidence on the priorities of the seizure community and those supporting them (eg, ref 101). To shape the project’s design and determine its perceived importance, a patient and public involvement (PPI) event for nine service users and their informal carers was completed. A similar exercise was completed with leading clinicians from seven of England’s ambulance services. Both groups were supportive of the project idea and provided feedback on the project’s draft design. When asked to rate its importance on a scale of 1–10, seven service user pairs scored it as 10 ‘Extremely important’.

Services users will be actively involved in the project’s completion. Service users are present in both the research team and in the groups advising and overseeing it. Coinvestigator JW is a service user herself with experience in ambulance care. Epilepsy Action, the largest seizure user organisation in the UK, is also a coinvestigator. A PPI group of 20 user representatives will contribute as research peers, advising the investigators on recruitment and reviewing study conclusions, implications for practice and recommendations. Four user representatives will also be on RADOSS Study Steering Committee (SSC).

All user representatives will be supported by Epilepsy Action who have an active PPI scheme and reimbursed for travel and their time according to guidance.102 Representatives will be recruited from a range of user groups.

**Ethics and dissemination**

Monitoring by an independent SSC will help to ensure the rights, safety and well-being of participants are the most important considerations. Compliance with the principles of Good Clinical Practice and scientific integrity will be managed by the study management team through regular and ad hoc meetings. YAS will be the sponsor. AJN and JMD are coinvestigators. WP1a and WP2 will use completely anonymised data from CUREd+. Access will be sought from the Centre for Urgent and Emergency Care Research Data Release Committee. CUREd+ has generic database ethical approval (307353) and Confidentiality Advisory Group approval (22/CAG/0019). With strict controls, WP1a and WP2’s work will be completed under these. WP1b and WP3 have received ethical approval from the University of Liverpool Central Research Ethics Committee D (11450). Only persons providing informed consent will participate.

We shall engage in a proactive dissemination and knowledge mobilisation strategy. It is specifically addressed by WP4 described in section Work package 4.

All requests for data sharing should be submitted to the corresponding author for consideration. Access to anonymised data may be granted following review.

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**Acknowledgements**

The authors would like to express their appreciation for the contributions from people who have experienced a suspected seizure and their family members and friends who will participate in this study or who have contributed data to CUREd+. They would also like to thank the members of the Study Steering Committee: Professor Niro Siriwardena (chair), Mrs Jayne Burton, Mrs Carol Jackson, Mr Graham Jackson, Ms Kim Kirby, Dr Benjamin Thomas and Professor Arjune Sen. The study sponsor is the Yorkshire Ambulance Service (yas. research@nhs.net) and they would like to thank Dr Fiona Bell and Ms Kelly Hird for their support with all aspects of the project set-up.

**Contributors**

AJN and JMD conceived the study and designed the study together with AGM, SMM, LJdB, MR, RF, JW, AF, RMJ, RMS and RC. LJdB planned the statistical analysis. AJN wrote the manuscript, with revisions being made by JMD, AGM, SMM, LJdB, MR, RF, JW, AF, RMJ, RMS and RC. All authors read and approved the final manuscript.

**Funding**

This project is funded by the National Institute for Health Research (NIHR) Research for Patient Benefit (RfPB) programme (reference NIHR203530).

**Disclaimer**

The views and opinions expressed herein are those of the authors and do not necessarily reflect those of the University of Liverpool, the RfPB programme, the NIHR, the NHS or the Department of Health and Social Care.

**Competing interests**

None declared.

**Patient and public involvement**

Patients and/or the public were involved in the design, or conduct, or reporting, or dissemination plans of this research. Refer to the Methods section for further details.

**Patient consent for publication**

Not applicable.

**Provenance and peer review**

Not commissioned; peer reviewed for ethical and funding approval prior to submission.

**Supplemental material**

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104 Baldwin T, Yorkshire ambulance service conveyance rates AMPDS 12, 2019.


SUPPLEMENTARY FILE 1-

FIGURE S1.1  Conceptual framework of factors affecting ambulance clinicians’ conveyance decisions (Oosterwold et al. [1])

**Notes:** The framework does not assert importance or interplay between factors. It does, though, highlight the quality of evidence support their importance (‘A’ is highest). That clinicians can find it difficult to confidently identify cases suitable for non-conveyance is captured under ‘Experience and confidence of the emergency medical service [EMS] provider’ and ‘Educational background, competencies and skills’ in the figure. Reproduced with permission (5347751335679).
REFERENCES

## SUPPLEMENTARY FILE 2

### TABLE S2.1

Range of structured data items that can be available to ambulance clinicians whilst on scene and will be included in data extract for RADOSS index events

<table>
<thead>
<tr>
<th>Data Items</th>
<th>Ambulance data system recorded within</th>
</tr>
</thead>
<tbody>
<tr>
<td>• Source of call (999, 111, police, other)</td>
<td>CAD</td>
</tr>
<tr>
<td>• Relationship of the caller to the patient</td>
<td>CAD</td>
</tr>
<tr>
<td>• Incident Date (day of week and month will be used)</td>
<td>CAD</td>
</tr>
<tr>
<td>• Incident Time(s)</td>
<td>CAD/PCR</td>
</tr>
<tr>
<td>• Patient age</td>
<td>PCR</td>
</tr>
<tr>
<td>• Patient sex</td>
<td>CAD</td>
</tr>
<tr>
<td>• Patient ethnicity</td>
<td>CAD</td>
</tr>
<tr>
<td>• Chief Complaint</td>
<td>CAD</td>
</tr>
<tr>
<td>• Dispatch AMPD code and description</td>
<td>CAD</td>
</tr>
<tr>
<td>• Priority category for response time</td>
<td>CAD</td>
</tr>
<tr>
<td>• Location type (private residence/ public place/ place of work/education)</td>
<td>PCR</td>
</tr>
<tr>
<td>• Incident location (Index of Multiple Deprivation 2020 and subdomain decile scores (IMD2020) and Rural Urban Classification of incident location (RUC11CD) will be used)</td>
<td>CAD/PCR</td>
</tr>
<tr>
<td>• Presenting complaint</td>
<td>PCR</td>
</tr>
<tr>
<td>• Clinician grade</td>
<td>PCR</td>
</tr>
<tr>
<td>• Was patient transported</td>
<td>PCR</td>
</tr>
<tr>
<td>• Type of facility transported to (ED, UTC, etc.)</td>
<td>PCR</td>
</tr>
<tr>
<td>• Reason for non-transport</td>
<td>PCR</td>
</tr>
<tr>
<td>• Referral to other service - type</td>
<td>PCR</td>
</tr>
<tr>
<td>• Pre-alert to hospital</td>
<td>PCR</td>
</tr>
<tr>
<td>• Physiological observations (e.g., pulse, BP, respiratory rate, oxygen saturation, level of consciousness AVPU, GCS, NEWS, blood sugar, temperature,)</td>
<td>PCR</td>
</tr>
<tr>
<td>• Previous medical history reported</td>
<td>PCR</td>
</tr>
<tr>
<td>• Airway intervention – type</td>
<td>PCR</td>
</tr>
<tr>
<td>• Wound care provided</td>
<td>PCR</td>
</tr>
<tr>
<td>• ECG findings</td>
<td>PCR</td>
</tr>
<tr>
<td>• Cardiac or respiratory arrest present</td>
<td>PCR</td>
</tr>
<tr>
<td>• Cardiac or respiratory arrest outcome</td>
<td>PCR</td>
</tr>
<tr>
<td>• Advice provided (non-transported patients)</td>
<td>PCR</td>
</tr>
<tr>
<td>• Supplementary oxygen provided</td>
<td>PCR</td>
</tr>
<tr>
<td>• Current medications</td>
<td>PCR</td>
</tr>
<tr>
<td>• Drugs administered (name, dosage, route)</td>
<td>PCR</td>
</tr>
<tr>
<td>• Time arrived scene</td>
<td>PCR</td>
</tr>
<tr>
<td>• Time departed scene</td>
<td>PCR</td>
</tr>
</tbody>
</table>

Notes: CAD, computer aid dispatch; PCR, patient care record; a Professional standards mean clinicians document findings from ≥1 set of observations[1]; b The RADOSS team will only have access to anonymised data; For reasons of governance and cost, the data extract for the index events will not
include free-text data entered by clinicians on ePCRs; Like most services, the Yorkshire Ambulance Service use the Advanced Medical Priority System (AMPDS) to code reason for call. When a call is received regarding what is described as a suspected seizure then AMPDS Protocol 12 is activated. It prompts call handlers to ask standard questions about the presentation severity and the patient. The responses inform the specific subcode given to the call. One question asked is whether the patient has a diagnosis of epilepsy (i.e., "Is s/he an epileptic?"); A minority (~5%) [2] of suspected seizure calls are transfers from ‘111’ rather than ‘999’ calls. As these calls were not managed via AMPDS, a slightly different data-set is available for them (e.g, disposition code). It will be included in the RADOSS data-extract. These items will have been calculated based on LSOA codes by the central CURE’d team who have access to pseudoanonymised data.
REFERENCES


SUPPLEMENTARY FILE 3

Details on the size of available linked data set and anticipated number of target events

Twelve-months of Yorkshire Ambulance Service (YAS) data should be sufficient to satisfy the requirements. It should permit a linked data-set to be formed for 13,980 index events [1, 2]. This is after excluding ~15% of cases that might have unlinkable records, ~3% who have ‘opted-out’ of research use of their data,[3] and the 0.02% of cases where the person died ‘on-scene’. [2]

Of the index events, 5,720 index events should have been conveyed of whom 229 (4%) will have experienced death/UEC recontact within 3 days and 1,201 (21%) could have experienced an AA. The remaining 2,451 index events should have been managed by non-conveyance, of whom 343 (14%) will have experienced death/ recontact with the urgent and emergency care system (UEC).[4-6]

The estimate that 14% of events not conveyed to ED will lead to death and/or recontact with the UEC within 3 days is based on O’Cathian et al.’s UK study.[4] We estimated that 4% of cases conveyed to ED will result in death and/or UEC contact. This estimate was generated differently since there is no UK evidence on this. Specifically, we considered Tohira et al.’s [5] US study. It reported adverse event rates within 24 hours in all persons who had attended ED for any reason were 2-4 times lower than for those not conveyed. It was also necessary to factor in individuals accounting for several types of events. Based on evidence from Coster et al.[7] we divided O’Cathian’s [4] estimates by 2 to account for this possibility and then applied Tohira et al. ratios to O’Cathian’s figures.

The estimate that 21% of seizure cases attending ED would satisfy the AA definition is informed by Miles et al.[6].

Key information factored into sample size calculation using Riley et al.’s formulae for each of the models

1. Permit testing of ≥40 candidate predictor parameters
2. Assume 0.05 acceptable difference in apparent and adjusted R-squared
3. Assume 0.05 margin of error in estimation of intercept
4. In the absence of other information, conservatively assumed that $R^2_{CS}$ for each model would correspond to an $R^2_{Nagelkerke}$ of 0.15. [8]
REFERENCES


