"The Score Matters": Wide Variations in Predictive Performance of 18 Paediatric Track and Trigger Systems

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(2479 words)

What is already known:

- Paediatric early warning systems (PEWS) are widely used to detect deterioration in hospitalised children
- The component parameters, weighting frameworks and scoring thresholds vary between differing PEWS
- Of the numerous PEWS in the literature and clinical practice, only a minority have been previously evaluated for their predictive performance

What this study adds:

- There is wide variation in the performance of PEWS
- There are no clear defining features which characterise the best performing PEWS
- The choice of PEWS may be an important factor in improving outcome for deteriorating hospitalised children

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Abstract

Objective

To compare the predictive performance of 18 Paediatric Early Warning Systems (PEWS) in predicting critical deterioration.

Design

Retrospective case-controlled study. PEWS values were calculated from existing clinical data and the area under the receiver operator characteristic curve (AUROC) compared.

Setting

UK tertiary referral children's hospital.

Patients

Patients without a 'do not attempt resuscitation' order admitted between 1st January 2011 and 31st December 2012. All patients on paediatric wards who suffered a critical deterioration event were designated 'cases' and matched with a control closest in age who was present on the same ward at the same time.

Main outcome measures

Respiratory and/or cardiac arrest, unplanned transfer to paediatric intensive care and/or unexpected death.

Results

Twelve 'scoring' and 6 'trigger' systems were suitable for comparative analysis. 297 case events in 224 patients were available for analysis. 244 control patients were identified for the 311 events.

Three PEWS demonstrated better overall predictive performance with an AUROC of 0.87 or greater. Comparing each system to the highest performing PEWS with Bonferroni's correction for multiple comparisons resulted in statistically significant differences for 13 systems. Trigger systems performed worse than scoring systems, occupying the 6 lowest places in the AUROC rankings.

Conclusion

There is considerable variation in the performance of published PEWS and as such the choice of PEWS has the potential to be clinically important. Trigger based systems performed poorly overall but it remains unclear what factors determine optimum performance. More complex systems did not necessarily demonstrate improved performance.

INTRODUCTION

Timely detection of evolving critical illness makes it easier to treat. Paediatric early warning systems (PEWS) should alert staff to deteriorating children and accelerate access to appropriate intervention. [1] Despite weak evidence [2,3] they are widely recommended.[4-8] In 2013 85% of UK centres caring for children were using a PEWS.[9]

Early warning systems are either 'score' or 'trigger'-based. Score-based systems assign values to vital signs (or other parameters), describing the variance from normal. These component values are then combined into an overall score. Higher scores should indicate reduced physiological reserve and prompt an escalating series of actions, culminating in senior clinician or rapid response team (RRT) review. The simpler 'trigger'-based systems contain thresholds for parameters without combining into an overall score. Again actions such as RRT review are often mandated. Scoring systems provide a more continuous description of the degree of abnormality in the child's physiological state compared to binary 'all or nothing' trigger systems.

The logic of standardised risk assessment is compelling, but the majority of PEWS have been developed using expert opinion alone. Comparative data are lacking on the relative performance of the 31 different published PEWS. Only a minority of these (14) have undergone *any* assessment of predictive validity.[1,10] Only one study compared the performance of multiple (3) scores.[11] Comparisons across studies are confounded by variance in the setting, methodologies, and outcomes described.[2]

Some might argue that the lack of validation or performance data is a secondary issue since the implementation of *any* system is the most important step. A system provides a structure for communication and builds consideration of risk of deterioration into daily practice. The alternative view is that the validity and calibration of any score are essential for utility. A score consistently providing false alerts while missing critical deteriorations elsewhere carries potential for harm by triaging resources incorrectly and increasing response times through 'alarm fatigue'.[12] Systems

have to balance specificity and sensitivity and so the precision of the thresholds included may be crucial.

We undertook a study comparing the performance of 18 PEWS in predicting critical deterioration in a UK tertiary referral children's hospital. Our null hypothesis was that the scores would show equivalent areas under the receiver operating characteristic curves.

METHODS

Evaluation of predictive validity

We undertook a retrospective case-control study of patients below 19 years of age without a 'do no attempt resuscitation' order who were admitted to our tertiary specialist children's hospital between 1st January 2011 and 31st December 2012. All patients who suffered a respiratory and/or cardiac arrest, unplanned transfer to PICU and/or unexpected death on the ward were designated 'cases'. They were identified from local data collected for the Paediatric Intensive Care Audit Network (PICANet) database,[13] the hospital resuscitation database and cross-referenced against intensive care admission records. Case patients present on the ward for less than 2 hours before the event were excluded as this was considered the minimum time for the child to be assessed, clinical signs recorded and action to be taken.

Case patients were each matched with a single control, present on the same ward at the same time. Wards were considered a proxy match for diagnostic speciality. The child closest in age to the case patient was identified. To ensure at least one set of observations could be extracted, control patients present on the ward for less than 24 hours were excluded, with the exception of wards classified as providing short stay/day case care where the threshold was 4 hours. Patients previously entered into the study were eligible to act as a control provided they did not suffer a critical deterioration event within the following 48 hours. If healthcare records were unavailable or the vital sign record was missing, the patient was excluded and a new control was sought using the same procedure.

Data extraction

Clinical data were extracted from the healthcare record of case patients for a period of 48 hours before the critical deterioration event. The final hour of data before the deterioration event in the case patient was excluded to establish if the PEWS could identify critical deterioration with at least one hour's notice. Data from controls was extracted for the same 47-hour period. Data were extracted by a single researcher (SC) using a standardised pro-forma. Vital signs were extracted as continuous variables. Respiratory effort was assessed retrospectively as mild, moderate or severe using standardised criteria.[14] Dichotomous variables were assessed using criteria in supplemental data table 1.

At the time of the study standard protocols were in place for recording and documenting vital signs, which nurses were informed of at induction and yearly intervals thereafter. The protocol mandated recording of a full set of vital signs within 2 hours of the start of the 12-hour shift. Elevated PEWS scores required repeat vital sign recording after 30 minutes. On-going frequency of recording was at the discretion of the bedside nurse.

Identification of Paediatric Early Warning Systems

Paediatric early warning systems were identified through our recent systematic review.[2] We excluded *a priori* PEWS where vital signs were assessed subjectively or against individual patient baseline values. Components of the remaining systems were reviewed to confirm that they could be extracted from the healthcare records. Criteria for data extraction were developed for included parameters (supplemental data table 1) together with the weighting framework for scoring systems. Minor inconsistencies such as overlapping age bandings were modified in a consistent manner to facilitate score calculation (Supplemental data table 2). Our hospital's local unpublished PEWS (children's early warning score [CEWS]) was also included (Supplemental data table 3).

Paediatric early warning system score calculation

Data were electronically checked for internal consistency and manually checked for accuracy. Inconsistencies were resolved by reviewing the data extraction proforma and healthcare records.

A recording of one or more vital signs was considered an observation data set. The PEWS value for each system was calculated for each observation data set. Missing observations were presumed to be normal (score 0), consistent with clinical practice and the methodology of previous studies [11,15,16]

Data analysis:

Analysis was performed using SPSS and r (www.cran.r-project.org). The maximum observed value for each PEWS for each patient in the 47 hours before the event was used in the comparison. Characteristics of cases and controls were compared with the Mann Whitney U-test for continuous variables and Chi-squared for categorical variables. Sensitivity, specificity, positive predictive value, negative predictive value, likelihood ratio positive test and likelihood ratio negative test were calculated. The area under receiver operator characteristic curves (AUROC) was calculated for each PEWS and compared with the best performing system using Delong's test for correlated curves.[17] Significance testing was adjusted for the multiple comparisons of AUROC with Bonferroni's correction, meaning p-values <0.0025 were considered significant.

The score that maximised sensitivity and specificity for each scoring system was identified as the optimal score.[20] The number of case and control patients who would be correctly and incorrectly identified at this threshold was calculated.

RESULTS

Characteristics of the identified paediatric early warning systems

Thirty-one PEWS were identified by the systematic review.[2] Seven contained parameters requiring subjective assessment, 6 required knowledge of the baseline vital signs and 1 inadequately described the component parameters: these were excluded. The remaining systems plus our local CEWS resulted in 18 PEWS. Systems with the same name were numbered in order of publication to distinguish between them (Table 1).

Twelve PEWS were 'scoring' and 6 were 'trigger' systems. The number of component parameters varied from 3 to 19. Some systems combined two or more variables within a single parameter, for example oxygen therapy and saturation values. Forty variables either alone or in combination were identified.

Vital signs were prominent. All 18 PEWS included heart and respiratory rate, 13 included oxygen saturation (72%) and 11 blood pressure (61%). Temperature was a component of only 7 systems (39%). Five weighting frameworks were identified across the 12 scoring systems, with 3 PEWS also incorporating additional points for risk factors. Differences between systems were often minor. The maximum scores varied from 7 to 32 (Table 1).

Patient characteristics

We identified 319 critical deterioration events. In 8 episodes the patient was present on the ward for less than two hours, leaving 311 eligible critical deterioration events in 237 patients. Fourteen case patient records were missing, leaving a case sample of 297 events in 224 patients. 244 control patients were identified for the 311 events.

In total 13551 observations sets were performed, 8360 on cases and 5191 on controls. The median number of observation sets per patient per day was 13 for cases and 6 for controls. Only 36.4% of

observation sets contained the 5 vital sign parameters and assessment of consciousness required for complete recording of the local PEWS.

Case patients were more likely to be female (56.3% vs 46.3%, p=0.009), have been admitted as an emergency (64.6% vs 39.2%, p=<0.01) and have a longer hospital stay (median 57.1 vs 35.9 days, p=<0.01). Mortality was also higher for case patients at 24 hours, 30 days and hospital discharge (p=<0.001). A summary of patient characteristics is shown in Table 2.

186 (62.6%) critical deterioration events were categorised as unplanned transfers to the PICU, 84 (28.3%) respiratory arrests and 27 (9.1%) cardiac arrests. Thirty-one patients remained on the ward after a cardiac or respiratory arrest. Six patients died before transfer to intensive care.

Predictive Performance

Three PEWS demonstrated better performance overall (Table 3). Comparing each system to the highest performing PEWS resulted in statistically significant differences for 13 systems. Overall trigger systems performed worse than scoring systems, occupying 6 of the lowest 7 places in the AUROC rankings.

Sensitivity, specificity, PPV, NPV and positive and negative likelihood ratio for the optimal score are given in Table 4. Values for trigger systems represent the breech of one or more trigger thresholds.

Trigger systems demonstrated better sensitivity (range 0.90-0.96) than scoring systems (range 0.46-0.83), but worse specificity (range 0.28 -0.56 versus 0.65 -0.91 respectively).

Our local PEWS performed modestly, ranked 10th overall. Comparison to the highest performing PEWS demonstrates the significantly worse predictive ability (Figure 1). At the optimal score the Cardiff and Vale PEWS would correctly identify 59 more deteriorating patients than our local PEWS, with only 4 additional false alerts.

Paediatric early warning systems demonstrated the ability to detect children at risk of critical deterioration a significant time before the event. Median time from optimal score [20] to event ranged from 17 hours (IQR 6.8-35.7) to 39.5 hours (IQR 17.4-46.6) for patients correctly identified by scoring systems. Longer times were demonstrated by trigger systems: 27.9 (IQR 13.7-42.4) to 39.8 hours (IQR 23.8-46.2), reflecting the increased sensitivity (Table 4).

DISCUSSION

The choice of PEWS is potentially important. Effective identification of 'at risk' children is crucial, but a poorly validated system may also erode staff confidence, waste valuable resources and overburden staff with false alerts. This study found that performance varied widely. Eight PEWS were good predictors, nine as useful and two poor.[33] Score-based systems consistently outperformed trigger systems. A larger number of parameters did not appear to improve performance, for instance the 2 lowest ranked systems had 16 and 14 parameters respectively compared to 8 parameters of the highest ranked system.

The Cardiff and Vale PEWS, Bedside PEWS and Modified PEWS III performed better than the majority of scores but with no significant different between them There were no obvious reasons why these systems outperformed the others. All three systems included heart and respiratory rate, oxygen saturation and blood pressure.

At the optimal score, scoring systems demonstrated poorer sensitivity, but superior specificity than trigger systems, which may reduce false alerts and build clinician confidence. Lowering the scoring thresholds improves sensitivity, creating additional opportunities to intervene and potentially improve outcome.[34] The ability to select the threshold that balances sensitivity and specificity most appropriate to the local environment gives scoring systems some advantages. However they are more complex to use, carrying the risk of inaccurate calculation [35,36] and inappropriate response [37,38].

The current local PEWS performed only modestly, despite being developed by local clinicians, using local data and expertise. It was considerably out-performed by systems externally validated in similar and differing populations. We have no reason to believe our situation is unique. It is likely that many other locally-developed unvalidated PEWS would demonstrate similar performance if evaluated rigorously. We are considering changing to the Bedside PEWS as it has now been evaluated in similar populations, is subject to an international multi-centre trial [10] and demonstrated equivalent performance with the top-ranked PEWS. This may facilitate further collaborative research in the future.

All PEWS demonstrated the ability to identify deteriorating children a number of hours before the event. Median hours from optimal score to critical deterioration event varied from 17.0 to 39.5 hours for scoring systems and 27.9 to 39.8 hours for trigger systems. This is longer than previous study findings for comparable scoring thresholds.[16] Both scoring and trigger systems can act as an important 'early warning' to front line staff of ward-based children at risk of critical deterioration, but require appropriate escalation and intervention by healthcare staff. Studies have identified that this may not always be achieved in practice.[6,39,40]

Limitations

Values for PEWS were calculated retrospectively from data extracted by a single researcher who was not blinded to the patient's outcome. Although standardised criteria were applied there was no other verification of data and accuracy of documented vital sign values and other observations could not be tested. Administration of a fluid bolus could not be reliably extracted affecting 3 PEWS.[15,19,27]

Data sets were frequently incomplete. Missing values were assumed to be 'normal' (score 0), but a recent study identified a greater proportion of incomplete data sets were associated with 'critical' (elevated) score compared to complete data sets.[36]. Incomplete vital sign recording remains an problem in clinical practice [21,41] and may under-estimate PEWS performance.

The study was conducted in a tertiary specialist children's hospital without an emergency department. Results may not be generalisable to children in other settings. Different results may also be seen for different outcomes and combinations of outcomes. Greater standardisation of reporting and consensus on pragmatic measures to evaluate PEWS and other similar interventions would facilitate meaningful comparison and collaborative research.[42]

CONCLUSION

The choice of PEWS may be important. Trigger based systems performed poorly overall but it remains unclear what factors determine optimum performance. More complex systems did not necessarily demonstrate improved performance. Variation in performance has important implications for effective identification of children 'at risk', staff confidence in the system and effective use of resources.

It is likely that many other hospitals have developed their own systems without rigorous evaluation of their validity.[43] The high and increasing number of both published and unpublished PEWS raises concerns that paediatrics may be following a similar path to that of adult track and trigger systems, with multiple poorly validated systems with unknown predictive power. This may explain why studies of PEWS and rapid response systems have so far failed to deliver the expected benefits.

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Table 1: Key characteristics and parameters

System																						Additional risk factors	
												Pa	aran	nete	ers (sco	red	usin	ıg w	eigł	nting framework)	Score 1 for each unless otherwise	
																						indicated	
						١	/ital	sign	IS		Con	cern		Other parameters							Other parameters		논
Name, first citation	Score or trigger	Maximum score	Age ranges	Parameters (n)	Heart rate	Respiratory rate	Oxygen saturation	Systolic BP	Capillary refill time	Temperature	Staff concern	Parent concern	Respiratory	Behaviour	Cardiovascular	Consciousness	Seizure	Respiratory distress	Airway threat	Oxygen therapy			Weighting framewo
Bedside PEWS[18]	S	26	5	7	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark									\checkmark		\checkmark			0,1,2,4
Bristol PEW tool [19]	Т	13	1	14	√2	✓	√1		✓		✓					√	√			✓	Apnoea ±bradycardia; DKA; clinically tiring or complete airway obstruction; hyperkalaemia; nebulised adrenaline; signs of shock (e.g. poor perfusion, ± low BP); suspected meningococcus		Trigger
Cardiff and Vale PEWS [20]	S	8	5	8	√	√	√	√			√					√		\checkmark	√	√			0, 1
Children's Early Warning System	S	21	4	6	√	√	√	√		√						√							0,1,2,3, 4
Children's Early Warning Tool [21]	S	24	4	9	√	√	√	√	√	✓						√		\checkmark		√			0,1,2,3
ITAT [22]	S	8	5	4	\checkmark	\checkmark	\checkmark			\checkmark													0,1,2
MET activation criteria I [23]	Т	9	5	9	√	\checkmark	\checkmark	√			√					✓	✓	\checkmark	√		Cardiac/respiratory arrest; apnoea or cyanosis		Trigger
MET activation criteria II [24]	Т	9	5	9	\checkmark	√	√	√			\checkmark	~				√	✓	\checkmark	√		Cardiac/respiratory arrest; apnoea or cyanosis		Trigger

System																						Additional risk factors	
												Pa	aran	net	ers	(sco	red	usi	ng w	/eig	hting framework)	Score 1 for each unless otherwise	
																						indicated	
						V	/ital	sigr	าร		Con	cern									Other parameters		¥
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	Scor	Max	Age	Para	Hea	Resp	0xy	Syst	Capi	Tem	Stafi	Pare	Resp	Behä	Carc	Con	Seiz	Resp	Airw	ŇXO			Š
Modified Bristol	Т	15	5	16	√2	\checkmark	\checkmark		\checkmark		\checkmark					\checkmark	\checkmark	\checkmark			Apnoea ±bradycardia; Clinically tiring or		Trigger
PEWS [25]																					complete airway obstruction; Hyperkalaemia;		
																					Marked increased work of breathing;		
																					Nebulised adrenaline (or no improvement);		
																					pH <7.2, Poor perfusion, ± low BP, large		
																					central/peripheral temp gradient; Unresolved		
																					pain on current anagesia therapy		
Modified PEWS I [26]	S	9	1	3	\checkmark	\checkmark			\checkmark				\checkmark	\checkmark	\checkmark					\checkmark			0,1,2,3
Modified PEWS II [27]	S	26	5	18	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark						\checkmark		\checkmark		\checkmark	CVL; IV bolus fluid or blood product within	Abnormal airway or positive pressure	0,1,2
																					past 4 hours	ventilation; Active acquired/congenital	
																						heart disease or history of heart surgery;	
																						Home oxygen; Pre/post any transplant;	
																						Gastrostomy or jejunostomy tube;	
																						Previous ICU admission; Severe	
																						developmental, neurological or	
																						neuromuscular disease	
Modified PEWS III	S	28	5	8	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark								\checkmark		\checkmark			
[28]																							

System																	Additional risk factors						
												Pa	aran	nete	rs (scoi	red	usin	ng w	eigł	nting framework)	Score 1 for each unless otherwise	
																						indicated	
						V	ital	sign	IS		Con	cern									Other parameters		k
Name, first citation	Score or trigger	Maximum score	Age ranges	Parameters (n)	Heart rate	Respiratory rate	Oxygen saturation	Systolic BP	Capillary refill time	Temperature	Staff concern	Parent concern	Respiratory	Behaviour	Cardiovascular	Consciousness	Seizure	Respiratory distress	Airway threat	Oxygen therapy			Weighting framewo
NHSI PEWS [29]	S	7	4	7	\checkmark	\checkmark					✓	√				\checkmark		\checkmark		\checkmark	Stridor or apnoea		0,1
PEW score I [30]	S	10	1	4	\checkmark	√			\checkmark			\checkmark	✓	√ \	1					√			0,1,2,3
PEW score II [16]	S	13	1	4	\checkmark	√			\checkmark				✓	√ \	1					√		15 minute nebulisers or vomiting post-op	0,1,2,3
PEW system score [15]	S	32	5	19	~	✓	1	~	~	~						✓				~	bolus fluid; pulses	>3 medical specialities involved in care; abnormal airway (not tracheostomy); CVL; gastrostomy; home oxygen; medication score; previous admission to ICU; severe cerebal palsy; transplant recipient	0,1,2,3
PMET triggers [31]	Т	7	5	8	\checkmark	\checkmark	√1	\checkmark			✓	√				\checkmark	\checkmark			\checkmark			Trigger
THCS MET calling criteria [32]	Т	7	1	7	\checkmark	V	A	\checkmark	√		√					\checkmark	\checkmark	\checkmark	√	√	Poor peripheral pulses, mottled extremities		Trigger

<u>Key:</u>

Indicators combined within a single parameter are presented in coloured text / 🗸 All studies are single centre unless otherwise stated. Overall risk of bias: L: Low; H: High Q: Qualitative or

quality improvement study and therefore risk of bias not assessed.

¹Separate parameters for children with and without cyanotic heart disease

² following one bolus of 10mls/kg fluid

Abbreviations: BP: Blood pressure; CVL: Central venous line; DKA: Diabetic ketoacidosis; ICU: Intensive Care Unit; ITAT: Inpatient triage, assessment and treatment score; IV: Intravenous; MET: Medical Emergency Team; NHSI: NHS Institute; PEW: Paediatric/Pediatric Early Warning; PEWS: Paediatric/Pediatric Early Warning System; PMET: Pediatric Medical Emergency Team; SVT: Super ventricular tachycardia; THSC: Toronto Hospital for Sick Children;

	Cases (n=297)	Controls (n=311)	P value
	n (%)	n (%)	
Gender			
Male	130 (43.8%)	167 (53.7)	0.018ª
Female	167 (56.3)	144 (46.3)	
Age			
0-<6 months	70 (23.6)	66 (21.2)	0.910 ^b
6 mo-<1y	41 (13.8)	47 (15.1)	
1y-<4y	87 (29.3)	94 (30.2)	
4y-<10y	49 (16.5)	55 (17.7)	
10y-<19y	50 (16.8)	49 (15.8)	
Gestation below 37 weeks	60 (20.1)	48 (15.4)	0.152ª
Weight, median, (interquartile			
range)	10.4kg (1.71-87.00)	11.1kg (2.10-94.20)	0.668 ^b
Previous same hospital admission			
0	150 (50.5)	145 (46.6)	0.946 ^b
1-5	66 (22.2)	92 (29.6)	
6-10	29 (9.8)	27 (8.7)	
11 – 20	20 (6.7)	26 (8.4)	
21 – 50	25 (8.4)	16 (5.2)	
>50	7 (2.4)	5 (1.6)	
Previous ICU admission (before this			
admission)			
0	247 (83.1)	276 (88.7)	0.061 ^b
1	32 (10.8)	20 (6.4)	
2	15 (5.1)	4 (1.3)	
3 - 5	1 (0.3)	5 (1.6)	
>5	2 (0.7)	6 (1.9)	
Previous PICU admission this episode			
0	185 (62.3)	238 (76.5)	<0.01 ^b
1	75 (25.2)	55 (17.7)	
2	17 (5.7)	14 (4.5)	
3 - 5	14 (4.7)	4 (1.3)	
>5	6 (2.0)	0 (0.0)	
Admitting speciality			
Medical	186 (62.6)	205 (65.9)	0.19 ^a
Surgical	57 (19.2)	66 (21.2)	
Intensive Care	54 (18.2)	40 (12.9)	

Table 2: Patient characteristics (each patient episode)

	Cases (n=297)	Controls (n=311)	P value
	n (%)	n (%)	
Type of admission			
Elective	105 (35.4)	189 (60.8)	<0.001 ^a
Emergency	192 (64.6)	122 (39.2)	
Speciality at event			
Medical	228 (76.8)	237 (76.2)	1.0ª
Surgical	69 (23.2)	74 (23.8)	
Critical deterioration event			
PICU transfer	186 (62.6)	0	N/A
Respiratory arrest	84 (28.3)	0	
Cardiac arrests	27 (9.1)	0	
Death on ward	0 (0)	0	
Reason for event			
Respiratory	176 (59.3)	0	N/A
Cardiovascular	67 (22.6)	0	
Neurological	38 (12.8)	0	
Other	16 (5.4)	0	
Length of stay in days, median,			
(interquartile range)	57.1 (21.0 – 122.0)	35.9 (12.8 – 89.4)	0.001 ^b
Length of hospital stay			
< 1 day	2 (0.7)	6 (1.9)	0.021ª
< 7 days	22 (7.4)	39 (12.5)	
< 30 days	74 (24.9)	91 (29.3)	
≥ 30 days	199 (670)	175 (56.3)	
Outcome			
Alive at 24 hours	279 (93.9)	311 (100%)	<0.001ª
Alive at 30 days	246 (82.8)	308 (99.0)	<0.001ª
Alive at discharge	220 (74.1)	301 (96.8)	<0.001ª

Key: ^aChi-squared; ^bMann-Whitney

Table 3: Comparative performance

Scoring systems	AUCROC (95% CI)	z-score	p-value
Cardiff and Vale PEWS	0.89 (0.86-0.91)	N/A	N/A
Bedside PEWS	0.88 (0.85-0.91)	0.72	0.47
Modified PEWS III	0.87 (0.85-0.90)	1.58	0.11
CEWT	0.85 (0.82-0.88)	3.21	0.001
Modified PEWS II	0.85 (0.82-0.88)	2.87	0.004
PEWS I	0.83 (0.80-0.86)	4.06	<0.001
NHSI PEWS	0.82 (0.79 - 0.86)	4.52	<0.001
PEWS system score	0.82 (0.78-0.85)	4.42	<0.001
PEWS II	0.79 (0.75-0.82)	6.00	<0.001
CEWS	0.79 (0.75-0.82)	7.12	<0.001
ITAT score	0.77 (0.74-0.81)	7.12	<0.001
Modified PEWS I	0.74 (0.70-0.78)	8.06	<0.001
Trigger systems			
THSC MET calling criteria	0.73 (0.69-0.77)	9.31	<0.001
MET activation criteria I	0.71 (0.70-0.75)	10.70	<0.001
MET activation criteria II	0.71 (0.70-0.75)	10.70	<0.001
PMET triggers I	0.71 (0.67 – 0.75)	10.82	<0.001
Modified Bristol PEWS	0.62 (0.58-0.67)	16.01	<0.001
Bristol PEWS	0.62 (0.58-0.67)	16.01	<0.001

Performance was assessed by calculation of the AUROC. Systems were then ranked and performance was compared to the highest ranked PEWS (Cardiff and Vale PEWS) using the Delong's test for correlated curves. z-scores represent comparison of mean values. Significance testing was adjusted for the multiple comparisons of AUROC with Bonferroni's correction, meaning p-values <0.0025 were considered significant.

<u>Abbreviations:</u> AUROC: Area under the receiver operator characteristic curve; ITAT: Inpatient triage, assessment and treatment score; MET: Medical Emergency Team; NHSI: NHS Institute; PEW: Paediatric/Pediatric Early Warning; PEWS: Paediatric/Pediatric Early Warning System; PMET: Pediatric Medical Emergency Team; THSC: Toronto Hospital for Sick Children

Table 4: Performance at	optima	score
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PEWS (AUROC	Optimal	Case patients		Control p	atients	Sensitivity	Specificity	PPV	NPV	LR +ve	LR –ve	Median hours
rank)	score/	(n=297)		(n=311)		(95% CI)	(95% CI)	to event				
	maximum score	Score at or above threshold	Score below threshold (FN)	Score at or above threshold	Score below threshold (TN)							(inter- quartile range)
Scoring systems		(ТР)		(FP)								
Scoring systems												
Cardiff and Vale	3/8	238	59	44	267	0.80	0.86	0.84	0.82	5.66	0.23	26.60
PEWS (1)						(0.75-0.84)	(0.81-0.89)	(0.80-0.88)	(0.77-0.86)	(4.28-7.49)	(0.18-0.29)	(11.57-42.19)
Bedside PEWS	6/26	215	82	35	276	0.72	0.89	0.860	0.77	6.43	0.311	26.25
(2)						(0.67-0.77)	(0.85-0.92)	(0.81-0.90)	(0.72-0.81)	(4.67-8.86)	(0.26-0.37)	(13.94-43.29)
Modified PEWS	7/28	204	93	28	283	0.69	0.91	0.879	0.75	7.63	0.34	21.61
III (3)						(0.63-0.74)	(0.87-0.94)	(0.83-0.92)	(0.71-0.80)	(5.31-10.95)	(0.29-0.41)	(12.42-40.10)
Modified PEWS	6/26	228	69	63	248	0.83	0.71	0.731	0.81	2.85	0.25	36.57
II (4)						(0.78-0.87)	(0.66-0.76)	(0.68-0.78)	(0.76-0.85)	(2.38-3.42)	(0.19-0.32)	(16.57-46.00)
CEWT (4)	4/24	245	52	90	221	0.77	0.80	0.784	0.78	3.79	0.29	37.66
						(0.72-0.81)	(0.75-0.84)	(0.73-0.83)	(0.73-0.83)	(3.01-4.77)	(0.24-0.36)	(22.39-44.74)
PEWS I (6)	3/10	247	50	99	212	0.83	0.68	0.714	0.81	2.61	0.25	24.00
						(0.78-0.87)	(0.63-0.73)	(0.66-0.76)	(0.76-0.85)	(2.20-3.10)	(0.19-0.32)	(11.23 –44.13)
NHSI PEWS (7)	2/7	247	50	108	203	0.83	0.65	0.696	0.80	2.40	0.26	29.90
						(0.78-0.87)	(0.60-0.71)	(0.65-0.74)	(0.75-085)	(2.04-2.81)	(0.20-0.33)	(14.57-43.63)
PEWS system	9/32	207	90	78	233	0.70	0.75	0.726	0.72	2.78	0.40	39.50
score (7)						(0.64-0.75)	(0.70-0.80)	(0.67-0.78)	(0.70-0.77)	(2.26-3.42)	(0.34-0.48)	(17.43-46.57)
PEWS II (9)	4/13	181	116	50	261	0.61	0.84	0.784	0.69	3.79	0.47	26.00
						(0.55-0.67)	(0.79-0.88)	(0.72-0.83)	(0.64-0.74)	(2.89-4.96)	(0.40-0.54)	(11.75-41.58)

PEWS (AUROC	Optimal	Case patie	ents	Control p	atients	Sensitivity	Specificity	PPV	NPV	LR +ve	LR –ve	Median hours
rank)	score/	(n=297)		(n=311)		(95% CI)	to event					
	maximum	Score at	Score below	Score at	Score below							(inter-
	score	or above	threshold	or above	threshold							quartile
		threshold	(FN)	threshold	(TN)							range)
	4/24	(TP)	110	(FP)	262	0.00	0.05	0.700	0.00	2.01	0.47	21.05
CEWS (9)	4/21	179	118	48	263	0.60	0.85	0.789	0.69	3.91	0.47	21.05
						(0.54-0.66)	(0.80-0.88)	(0.73-0.84)	(0.64-0.74)	(2.96-5.15)	(0.41-0.54)	(10.38-40.12)
ITAT score (11)	3/8	202	95	82	229	0.68	0.74	0.711	0.71	2.58	0.43	28.95
						(0.62-0.73)	(0.68-0.78)	(0.65-0.76)	(0.65-0.76)	(2.11-3.16)	(0.37-0.51)	(14.70-43.96)
Modified PEWS I	4/9	135	162	31	280	0.46	0.90	0.813	0.63	4.56	0.61	17.00
(12)						(0.40-0.51)	(0.86-0.93)	(0.74-0.87)	(0.59-0.68)	(3.19-6.51)	(0.55-0.67)	(6.75-35.68)
Trigger systems				-								
THSC MET	1 or	267	30	138	173	0.90	0.56	0.66	0.85	2.03	0.18	27.90
calling criteria	more					(0.86-0.93)	0.50-0.61)	(0.61-0.71)	(0.79-0.90)	(1.78-2.31)	(0.13-0.26)	(13.74-42.37)
(13)	triggers											
MET activation	1 or	276	21	158	153	0.93	0.49	0.64	0.88	1.83	0.14	33.87
criteria I (14)	more					(0.89-0.96)	(0.44-0.55)	(0.59-0.68)	(0.82-0.92)	(1.63-2.05)	(0.10-0.22)	(18.76-45.52)
	triggers											
MET activation	1 or	276	21	158	153	0.923	0.49	0.64	0.88	1.83	0.14	33.92
criteria II (14)	more					(0.89-0.96)	(0.44-0.55)	(0.59-0.68)	(0.82-0.92)	(1.63-2.05)	(0.10-0.22)	(18.76-45.52)
	triggers											
PMET triggers I	1 or	273	24	157	154	0.92	0.50	0.64	0.87	1.82	0.16	33.25
(14)	more					(0.88-0.95)	(0.44-0.55)	(0.59-0.68)	(0.80-0.68)	(1.62-2.04)	(0.11-0.24)	(16.90-45.42)
	triggers											
	0.000.0			1								

PEWS (AUROC	Optimal	Case patie	ents	Control pa	atients	Sensitivity	Specificity	PPV	NPV	LR +ve	LR –ve	Median hours
rank)	score/	(n=297)		(n=311)		(95% CI)	to event					
	maximum	Score at	Score below	Score at	Score below							(inter-
	score	or above	threshold	or above	threshold							quartile
		threshold	(FN)	threshold	(TN)							range)
		(1P)		(FP)								•
Modified Bristol	1 or	286	11	223	88	0.96	0.28	0.56	0.90	1.34	0.13	39.83
PEWS (17)	more					(0.93-0.98)	(0.23-0.34)	(0.52-0.61)	(0.81-0.94)	(1.25-1.45)	(0.07-0.24)	(23.82-46.25)
	triggers											
Bristol PEWS	1 or	285	12	223	88	0.96	0.28	0.56	0.88	1.34	0.14	39.73
(17)	more					(0.93-0.98)	(0.23-0.34)	(0.52-0.61)	(0.80-0.93)	(1.24-1.44)	(0.08-0.25)	(23.45-46.25)
	triggers											

The optimal score for trigger systems was determined as 1. The optimal score for scoring systems was determined as the cut-point which demonstrated the maximum value for the sum of the sensitivity and specificity, as described by Edwards et al.[20] As such, this differed between different scoring systems.

The hours to event was calculated as the number of hours between the case patient's first achieving the optimal score/trigger to the occurrence of the critical deterioration event.

Abbreviations:_AUROC:_Area under the receiver operator characteristic curve; CI: Confidence interval; FP: False positive; FN: False negative; ITAT: Inpatient triage, assessment and treatment score; LR +ve; Positive likelihood ratio; LR –ve: Negative likelihood ratio; MET: Medical Emergency Team; NHSI: NHS Institute; NPV: Negative predictive value; PEW: Paediatric/Pediatric Early Warning; PEWS: Paediatric/Pediatric Early Warning System; PMET: Pediatric Medical Emergency Team; PPV: Positive predictive value; THSC: Toronto Hospital for Sick Children; TP: True positive; TN: True negative