A 3-Staged Approach to Identifying Patients at Risk of Deterioration in Emergency Departments

Thomas Schmidt and Uffe Kock Wiil

The Maersk Mc-Kinney Moller Institute, University of Southern Denmark, Campusvej 55, 5230 Odense M, Denmark

Keywords: Health Informatics, Patient Monitoring, Emergency Departments, Predictive Modelling.

Abstract: The variety in patient demographics and admission reasons makes it challenging for Emergency Department clinicians to notice deterioration in patients. Recent research has found that up to 20% of non-critical patients deteriorate within the first 24 hours after admission. Unnoticed patient deterioration can lead to serious adverse events in a clinical setting where patient monitoring relies solely on manual observations of monitors at infrequent intervals. In this paper, we present a novel 3-Stage Patient Deterioration Warning System as a model to mitigate the risk of undetected deterioration while improving clinical alarm fatigue. This staged approach enables the monitoring of patients in levels of increasing descriptiveness based on multiple models of normality. The model is validated via related work, clinical observations, and patterns of patient data collected at a Danish Emergency Department bedside ward. The paper concludes with a presentation of plans for future implementation work.

1 INTRODUCTION

Roughly 20% of patients arriving at an emergency department (ED) with non-critical vital values, deteriorate within the first 24 hours (Henriksen et al., 2014). In line with the assumption that 3 to 6% of all deaths in hospitals are unexpected (Hayward and Hofer 2001; Zegers et al., 2009), this has spawned several attempts to prevent these situations by increasing formalization and automation of patient observations.

In this regard much scholarly effort has addressed the challenge of improving the predictive accuracy of Early Warning Systems (EWS) for detecting patient deterioration. Broadly speaking, these attempts can be classified as either improving the system for identifying deteriorating patients or seeking to automate the deterioration detection system to alleviate the cognitive and physical workload on clinicians. Permutations of the first aspect have been researched with regards to improving EWS in general wards (Mcgaughey et al., 2007), and in EDs (Geier et al., 2013). A recurring theme in this research is the inability to definitively determine exact vital sign thresholds and correlations to mark the initial stage of deterioration. Thus, most systems have poor quality of supporting evidence (Gao et al., 2007; Brabrand et al., 2010).

Most approaches do however note an effect of implementing a formalized EWS. This is in line with the second line of attempts, where the process of identifying patients at risk depends on collaboration and communication between multiple actors. This interplay has been coined as the "chain of survival" (Subbe and Welch 2013). This chain consists of: 1) high-quality recording of vital signs; 2) ability to recognize familiar patterns; 3) reporting of abnormality; and 4) a precise and prompt response. Of these four steps, this paper focuses primarily on steps 2 and 3, by introducing a model that adheres to the patients' clinical circumstances, and to provide a system design that enables higher accuracy, while supporting the cognitive models of clinicians.

Our work includes a field study of an Emergency Department, with the purpose of identifying collaborative and organizational causes for undetected patient deterioration. This field study is based on 13 participatory observation sessions of complete 8 hour shifts with different clinical groups. We conclude that future patient deterioration detection systems must convey information about patient state and trajectory ubiquitously throughout the department, and not just at bedside or in designated offices to overcome both temporal and spatial challenges. Additionally, as each clinical group has different areas of expertise and means of

 470 Schmidt T. and Will U.. A 3-Staged Approach to Identifying Patients at Risk of Deterioration in Emergency Departments. DOI: 10.5220/0005226004700477 In *Proceedings of the International Conference on Health Informatics* (HEALTHINF-2015), pages 470-477 ISBN: 978-989-758-068-0 Copyright © 2015 SCITEPRESS (Science and Technology Publications, Lda.) interaction, the deterioration warning system should differentiate patient state representation according to individual clinical groups (Schmidt & Wiil n.d.).

From the field study, we also identified that clinical observations and automated patient monitoring are challenged by the fact that certain groups of patients and individuals fall outside the population based model of normality currently in use at the ED. Several attempts to improve the predictive accuracy of EWS have already been made (see Section 3). However, our findings indicate that models should accommodate the individual traits of each patient. Yet, the clinical reality often proceeds at a pace that prohibits this from being attainable in a real time environment. Thus, we present a design that counters these challenges through a staged model which allows for a gradual progression of accuracy as the system familiarizes itself with each patient. We have named this system the 3-Stage Patient Deterioration Warning System (3-Stage PDWS).

The paper first describes the settings and structure of a Danish ED. We then describe related work in Section 3 to argue for our approach. We present our data collection approach and the study methodology which sets the stage for the design of the 3-Stage PDWS. As this is the main contribution of the paper, we conclude with a discussion of the challenges we face and a description of planned future work to address these.

2 THE SETTINGS

The ED capacities of the Danish healthcare sector have recently been restructured by merging multiple hospital entry points to a single point of entry. This meant closing emergency departments at minor hospitals and fusing the capacities of larger hospitals. All observations in this paper are based on a field study conducted at an ED at a large university hospital in Denmark. This ED is organized into a receiving ward, known as the Emergency Treatment Center (ETC) and a bedside ward; Center for Accelerated Patient admissions (CAP). The ETC handles both medical and surgical illnesses which can be identified and treated in a day. If the treatment period requires hospitalization, the patient will be admitted to the CAP. All patients arriving at the ED with anything but minor injuries will be triaged upon arrival. The variety of patients in EDs makes it difficult to define a single warning system to suit all patients (Windle and Williams 2009). At the ED of this study, the clinicians currently rely on

the ADAPT triage model (Lauritzen et al., 2009), which defines thresholds for each severity score and provides guidelines for how often registered observations are to be scheduled during the stay.

Vital signs monitoring in the ED utilize Philips IntelliVue MP30/50 monitors in a networked setup which enables clinicians to remotely monitor patients from ward offices. How much and at what frequency a patient is monitored depends on clinical judgment based on the patient's triage level.

A distinctive trait of EDs is that the clinicians plan treatment of patients based on their presented history and symptoms instead of a known diagnosis. So in a context where patient throughput is high and a large part of clinical observations are tacit and thus seldom transferred consistently between shifts, the need for a shared representation which captures a patient's state, trajectory, and clinically linked observations is a reality that to the best of our knowledge is not dealt with properly today.

During our field study we observed on multiple occasions nurses muting patient alarms without actually assessing the patient's state. This oversight of alarms was frequently based on assumptions about the patient, or the equipment's reliability. This is in line with similar causing factors for alarm fatigue such as a high number of false positives, usability issues, and faith in own knowledge (Sijs et al., 2006). The monitoring system issues alarms in stepwise degrees, and even though the most severe alarms still lead to increased levels of observation, inexperienced nurses may be affected by the overall tendency to dismiss non-critical alarms, and thus miss true adverse events in the long run.

As few EDs to our knowledge have the necessary staffing and budget to integrate the latest generation of automated patient monitoring, we believe that there is a need to identify ways of improving deterioration detection by utilizing existing equipment. This pragmatic approach should be of interest to EDs worldwide.

3 RELATED WORK

In our review of existing related work, we have focused on studies that concentrated on integration into a clinical reality: work that attempts to integrate prospective data, real-time analysis, and an assessment of clinical feasibility. From these criteria, the research contributed by the Oxford Biomedical Research Center (Tarassenko et al., 2006; Orphanidou et al., 2009) stands out. They investigate the applicability of latent variable models which merge multiple streams of patient vital values into a model built upon machine learning techniques, with the intention of providing an intuitive visualization of patient state and trajectory.

The plausibility of building individual models of normality has been investigated (Zhang et al., 2007). Although the specific angle in this study is unfit for a large scale real-time system, the research still conceptually shows the possibility of detecting patient deterioration from dynamically created models. In a study based on observational vital sign data, models of normality were built for a specific post-operative patient population based on three different metrics calculated from the vital sign distributions (Pimentel et al., 2013). In the same study, the authors also found that the majority of observed vital sign types varied substantially from submission to admission.

Priming a clinical warning system by performing risk stratification based on Electronic Health Record (EHR) information to determine which patients were in need of continuous monitoring offers several advantages (Hackmann et al., 2011). This vision has been elaborated upon by focusing on the challenges of doing time series analysis on streams of vital signs (Mao et al., 2011).

Although several contributions to this field have been made, most of the work has been done in parallel, and not in cooperation, with the targeted clinical context. Thus, we are motivated to conduct the planning and execution of this project with the intent of providing a solution that strives to fit into the entirety of the problem domain.

4 METHODOLOGY AND DATA GATHERING

This paper is part of larger action-oriented research project which involves a field study, workshops, and prototype-driven controlled experiments. As such, we follow an action-oriented research approach (Easterbrook et al., 2008). Consequently, we have participated in ED training courses, managed workshops, and helped plan new standard working procedures. In October 2013, we launched an ongoing automated gathering of vital sign data from patients admitted to the CAP. The registration of vital signs is approved by the Danish Data Protection Agency. Data is stored in a restricted access database in compliance with Danish legislation on privacy concerns.

The collected vital signs will be coupled with national Danish health registries to cluster all

patients using categorical data such as past illnesses from ICD-10 codes, initial triage level, gender, admission package, number of prescribed medications, age, and {7,30,90}-day outcome in a retrospective analysis. The dataset will be segmented into event and non-event subgroups based on the occurrences of heart failures, ICU transfers, and inhospital death. This retrospective dataset also forms the foundation for the training of the patient state models which we introduce in later sections.

4.1 Vital Sign Data Collection

Vital sign values are harvested from the Philips IntelliVue patient monitors through a HL7 export interface. From this we receive HL7 Unsolicited Observation Reporting messages with patient vital signs from each bed in 60 second intervals. These messages are parsed and stored in a VitalSigns database. The HL7 messages carries information about arterial blood oxygen saturation (SpO2) and Pulse Rate (PR) measured through pulse oximetry; Respiration Rate (RR); and Heart Rate (HR) measured using 3-lead electrocardiography; and mean, systolic, and diastolic blood pressure measured using a Non-invasive Blood Pressure (NBP) cuff. The actual types of vital signs registered for each individual patient depend on the level of criticality and overall mobility of the patient. Clinicians often adjust the frequency of NBP measurements to match the state of the patient, and consequently we register blood pressure measurements in intervals from five to sixty minutes. As pulse oximetry is the least obtrusive vital sign to monitor, SpO2 and PR are by far the most frequent observations in our dataset.

When a patient is received on the CAP ward, we asked the nurses to admit the patient to the Philips IntelliVue system by entering personal identification

Number of patients registered	5.023
Total number of registered aggregated vital signs	3.483.302
Mean age male patients in years	61.4 (SD 20.4)
Mean age female patients in	65.4
years	(SD 22.3)
Heart Rate registrations	1.972.106
Respiratory Rate registrations	1.925.849
Pulse Rate registrations	3.015.059
SpO2 registrations	2.996.736
Blood Pressure registrations	112.589

Table 1: Overview of vital sign registrations.

information such as name and social security number. This information was stored in a Patient database table and coupled with the vital signs.

Table 1 summarizes the collection of vital values in the period from October 2013 to August 2014. The number of vital signs registered for each patient varies from a single measurement up to several thousand.

5 DESIGNING THE 3-STAGE PDWS

5.1 Guiding Design Principles

Our design principles are influenced by related work, clinical observations, and our own assumptions regarding what seems feasible fitting into the settings of the particular ED from the field study.

The ongoing data collection has been preliminarily evaluated to probe for support of our assumption that a granular model of normality would be an appropriate approach for the system. As an example of group-based normality, Figure 1 depicts the distributions of 6.000 randomly sampled heart rates of patients assigned to either the Endocrinology (E) or the Neurology (N) speciality; mean heart rate for each sample is shown as vertical lines.

Although Figure 1 displays distinct differences, medical specialty as such is not a sufficiently accurate classification feature. E patients in the ED are often diabetics whereas N patients can have a wider range of diseases. Later investigations will include ICD-10 codes with highest mortality and coverage of admissions.

For patients these will be obtained through the



Figure 1: HR distribution for two specialties.

coupling of the VitalSigns database to the Danish national health registries. Thus, the specific model features are currently being selected through our cooperation with ED researchers and clinicians.

Figure 2 illustrates the key assumption that patients arriving at the ED are classifiable as unstable as they are in an imbalanced condition compared to their normal state of being.



As clinicians initiate the diagnosis and treatment process during hospitalization, the patient is assumed to gradually stabilize, as illustrated by box 1 in Figure 2. Some patients are expected to stabilize during treatment, which marks the period where the 3-Stage PDWS trains to detect deterioration from personalized stabilization, illustrated as box 2 in Figure 2. This individualized approach is expected to enable earlier realization of the cases where a patient departs from individual stability, illustrated as box 3 in Figure 2. Automated recognition of deterioration is expected to precede human clinical observation of the same deterioration, which is shown as box 4 in Figure 2. Even if the discriminatory ability of the system is poor, it might still provide warning of ongoing deterioration at an earlier stage as the automated system continuously evaluates the patients, whereas clinicians are confined to spot observations due to a busy schedule.

The patient trajectory assumption will be tested by analysing the vital signs time series data collected from patients to check for any significant progression of vital signs during the admission period of the patients. Similar investigations have been carried out and found that the initial trend of a patient's state is correlated to in-hospital death (Kellett et al. 2013).

5.2 Model Architecture

The conceptual model in Figure 3 depicts each of the three stages in our proposed system:



Figure 3. The 3-Stage PDWS model.

This model utilizes three levels of deterioration detection; λ_p for detecting deviation from population based training set, $\lambda_g \in \lambda_{GROUP}$ from a set of group based models, and λ_i for individual calibrated models.

Each stage in Figure 3 serves a particular purpose:

- 1. *Population-based Deterioration Detection:* Initially, the state of the patient is derived from a population-based deterioration model.
- 2. Group-based Deterioration Detection: when a patient is classified as belonging to a given group $g \in GROUP$, all received vital
- values from the patient will be assessed according to what is identified as normality for this group of patients.
- 3. Personalized Deterioration Detection: given the assumption that some patients have models of normality that differ from any group, the system will evaluate if the given patient seems to be in a stable deviation from the model of normality under which the patient is currently monitored. If so, the patient should be monitored according to an individual model.

Between the 1st and 2nd stage, patients are sought classified based on arrival parameters and from information from the patient's EHR. This classification will be based on a previous unsupervised clustering of patient parameters. In this step, we initially seek to select a couple of the most significant clusters to reduce the model complexity. The envisioned flow of stage selection and state decoding is illustrated in Figure 4, which shows the parallel deterioration detection, training of the individual model, and concurrent visualization of patient state.

Although we are still evaluating machine learning techniques, using Hidden Markov Models (HMM) as a modelling approach for patient state transitions is interesting because the properties and traits of HMMs resemble the clinical reality found in EDs. Namely, that clinicians monitor a set of vital sign observations from which they seek to deduce

the actual state of a patient. This is in line with the hidden state nature of HMMs (Rabiner, 1989). Although clinicians operate with a multitude of observation channels, the clinical assessment is essentially still a process of uncertainty and interpretation of the hidden, actual state of the patient. HMMs have been used to model clinical relevant situations such as real-time daily activity monitoring (Wei et al., 2011) and hepatitis C disease progression (Sweeting et al., 2010). Although (Sittig and Factor, 1990) investigated the development of a multi-state Kalman filter algorithm for patient monitoring, and (Ghassempour et al., 2014) proposed a method for clustering multivariate time series of both numerical and categorical features in healthcare, our survey of published research indicated that HMMs have not been investigated for modelling patients in an ED context.



Figure 4: Stage selection flow.

Through our observations of patients and discussions with clinicians, it is evident that it is difficult to quantify the state of a patient. Instead the trajectory of a patient is often mentioned as a noticeable registration by clinicians, which raises the question if dynamic changes in patients can be used to identify patients at risk (Kellett et al., 2013). Hence, Figure 5 conveys our proposal for the states of the HMM and its transition relationships. The hidden states of the HMM are the unknown actual state of the patient who can be either in one of the safe states, transition states, or unsafe states.

The decision threshold between transitioning from group-based normality to individual normality will be investigated using distance measures between the two HMMs (Lyngsø et al., 1999; Zeng et al., 2010) and will also be evaluated based upon an evaluation of $P(O|\lambda_i) > P(O|\lambda_{active})$.



Figure 5. Proposed states.

5.3 Model Validation

The 3-Stage PDWS model has currently only been conceptually validated by its composition from the body of existing published research, preliminary data analysis, and clinical observations from the field study, and in collaboration with nurses and physicians at the ED.

The exact number of clusters found through the investigation of patient characteristics, is still unknown. However, we intend to validate the clusters by relying on the judgment of experienced ED physicians who will review the similarity measures of each cluster.

The final model and its implementation needs validation in two dimensions: a retrospective evaluation of its accuracy in identifying patients at risk of deterioration, and in its ability to convey the patient state in a way that makes sense to different clinical professions.

Each stage of the PDWS will be assessed individually in the retrospective validation by its ability to accurately classify patients as The accuracy deteriorating. the of group classification will be reviewed through the ability to produce similar labels for unobserved patients as found by the physicians. In this regard we are interested in model accuracy and the ability to predict deterioration onset earlier than currently possible by the existing alarming thresholds defined by the ADAPT triage model (Lauritzen et al., 2009).

The clinical utility of our model will be assessed by comparing the misclassification rate of our system with the generic thresholds used at the specific ED in this study.

6 **DISCUSSION**

Automating patient deterioration detection can be

approached from multiple entry points. The first challenge is gathering the vital signs in an unobtrusive way that does not enforce a potentially unjustified sense of illness on the patient, and which does not hinder the workflow of clinicians or treatment trajectories of patients. Our approach is pragmatic in the sense that we seek to design and build a solution that utilizes the existing equipment at the ED. We have found that the clinicians are prone to not attach the most cumbersome sensors to patients who are scheduled for frequent tests outside the ward. Mobile monitoring technology would help overcome this obstacle, but is outside the scope of our current research approach.

Our approach to subgroup classification resembles that of (Zmiri et al., 2012), who investigated the feasibility of using decision trees and probabilistic algorithms for classification of patients into severity levels similar to the clinical triage classification. However, our intent is not to replace existing severity indices, but instead to improve the accuracy of vital sign monitoring by deploying increasingly specific thresholds.

Although the data we are currently collecting only consist of a few dimensions, we are challenged by commonly found problems such as variation in what vital signs are measured and occasional holes in the time series. This issue has been dealt with by replacing the missing values with either the last registered measurement or with the mean of the vital sign over the entire historical dataset (Mao et al., 2011). An alternative approach is to utilize Gaussian Processes which have proved useful in predicting the distribution of missing physiological data (Clifton et al., 2012). Additionally, we face a sizeable task in ensuring the validity of the vital signs by having to check that the data series can be linked to a given patient in a reliable manner. To ensure this we plan to couple our VitalSigns database with the EDs internal logistic system. This provides accurate information about which patients resided in each bed at a given point in time.

Modifying the harvesting and registration of vital sign data using existing equipment by asking clinicians to revise their standard working practices, has unsurprisingly proven difficult. The staff group as a whole recognizes the importance of registering vital values. But the clinical reality is such that if a system does not yield immediate and tangible benefits, the perception of added utility is generally low, causing the clinicians to abstain from integrating new admission procedures. In our case this is admitting patients to the Philips IntelliVue system, but we find that the concept of clinical utility and the interplay between healthcare organization and health information technology is a topic worthy of further exploration. This interplay relates to the notion of "meaningful use" (Blumenthal and Tavenner, 2010), and the SUMMIT method for modelling the meaningful use of an IT tool as a function of its overall utility (Haynes et al., 2013). It seems feasible to deploy this framework in the planned controlled experiments with nurses and physicians to structure the evaluation of how the 3-Stage PDWS represents and visualizes knowledge and information about patients. Other research has pointed out a clinical scepticism towards black-box expert systems. Consequently, the aim of our system is to support decision making rather than replace it.

Finally, while our initial approach will rely on datasets tightly coupled to the Danish healthcare system, the core model assumptions are applicable globally. Although specialty department features as proposed in Figure 1 may be of little meaning to other healthcare systems, we expect that the addition of selected ICD-10 features will provide both interesting insight for clinical researchers and practitioners. The challenges of patient variation is known to all EDs and thus we believe that the 3-Stage PDWS can be of use wherever it is possible to classify arriving patients.

Another aspect is the availability of vital sign data. Several patient monitors already support exporting vital sign data, and as such our solution is independent of particular equipment. As monitoring platforms are becoming increasingly unobtrusive, a wider spectrum of patients can be included in continuous monitoring. This expansion into a broader part of the patient population further justifies building more specific deterioration detection models.

7 FUTURE WORK AND CONCLUSION

The overall goal is to improve the detection of deteriorating patients by identifying the onset of adverse events earlier and to embed this detection ubiquitously into clinical practices by assuming a holistic approach to the integration of patient monitoring. If the system proves successful, we expect to see a reduction in patient mortality and increased clinical utility of the monitoring platform.

The intent of our research is to target the solution domain as a whole and not to focus on particular parts, e.g., providing a revolutionary real-time analysis model, conceiving new machine learning techniques, or developing new monitoring platforms. We expect to draw out more systemic findings which can support more depth-oriented research approaches.

The system is currently under development, as we have undertaken initial analysis of the vital signs and how to utilize these with machine learning techniques that are sensible to clinicians. Coupling of the collected vital signs with the national Danish health registries is planned for Q3 2014, and we expect the prototype to be ready for initial clinical controlled experiments by early 2015. Fine tuning of the predictive capabilities of the 3-Stage PDWS is planned for Q2 in 2015.



- Blumenthal, D. & Tavenner, M., 2010. The "Meaningful Use" Regulation for Electronic Health Records. *New England Journal of Medicine*, 363(6), pp.501–504.
- Brabrand, M. et al., 2010. Risk scoring systems for adults admitted to the emergency department: a systematic review. *Scandinavian journal of trauma, resuscitation and emergency medicine*, 18, p.8.
- Clifton, L. et al., 2012. Gaussian process regression in vital-sign early warning systems. 2012 Annual International Conference of the IEEE Engineering in Medicine and Biology Society, pp.6161–6164.
- Easterbrook, S. et al., 2008. Selecting empirical methods for software engineering research. In F. Shull, J. Singer, & D. I. K. Sjøberg, eds. *Guide to Advanced Empirical Software Engineering*. Springer London, pp. 285–311.
- Gao, H. et al., 2007. Systematic review and evaluation of physiological track and trigger warning systems for identifying at-risk patients on the ward. *Intensive care medicine*, 33(4), pp.667–79.
- Geier, F. et al., 2013. Severity illness scoring systems for early identification and prediction of in-hospital mortality in patients with suspected sepsis presenting to the emergency department. *Wiener klinische Wochenschrift*, 125(17-18), pp.508–515.
- Ghassempour, S., Girosi, F. & Maeder, A., 2014. Clustering multivariate time series using Hidden Markov Models. *International journal of environmental research and public health*, 11(3), pp.2741–63.
- Hackmann, G. et al., 2011. Toward a two-tier clinical warning system for hospitalized patients. In *AMIA Annual Symposium Proceedings*. pp. 511–9.
- Haynes, S.R., Winkler, T.E. & Ritter, F.E., 2013. Modeling Meaningful Use as Utility in Emergency Medical Services. 2013 IEEE International Conference on Healthcare Informatics, pp.455–460.
- Hayward, R.A. & Hofer, T.P., 2001. Estimating hospital deaths due to medical errors: preventability is in the

eye of the reviewer. *The Journal of the American Medical Association*, 286(4), pp.415–420.

- Henriksen, D.P., Brabrand, M. & Lassen, A.T., 2014. Prognosis and risk factors for deterioration in patients admitted to a medical emergency department. *PloS* one, 9(4), p.e94649.
- Kellett, J. et al., 2013. Changes and their prognostic implications in the abbreviated VitalPACTM Early Warning Score (ViEWS) after admission to hospital of 18,827 surgical patients. *Resuscitation*, 84(4), pp.471– 6.
- Lauritzen, M., Skriver, C. & Dahlin, J., 2009. Triage-Manual. , (juni). Available at: http://www.hillerodhospital.dk/NR/rdonlyres/D20F6C 68-ABB6-402D-B463-C7293185C372/0/Triagemaster.pdf.
- Lyngsø, R.B., Pedersen, C.N. & Nielsen, H., 1999. Metrics and similarity measures for hidden Markov models. In *Proc. Int. Conf. Intell. Syst. Mol. Biol.* pp. 178–86.
- Mao, Y., Chen, Y. & Hackmann, G., 2011. Medical Data Mining for Early Deterioration Warning in General Hospital Wards. In *Proceedings of the 2011 IEEE 11th International Conference on Data Mining Workshops*. IEEE Computer Society, pp. 1042–1049.
- Mcgaughey, J. et al., 2007. Outreach and Early Warning Systems (EWS) for the prevention of Intensive Care admission and death of critically ill adult patients on general hospital wards (Review). *Cochrane Database System Rev*, 3.
- Orphanidou, C. et al., 2009. Telemetry-based vital sign monitoring for ambulatory hospital patients. In Engineering in Medicine and Biology Society, 2009. EMBC 2009. Annual International Conference of the IEEE. pp. 4650–3.
- Pimentel, M.A.F. et al., 2013. Modelling physiological deterioration in post-operative patient vital-sign data. *Medical & biological engineering & computing*, 51(8), pp.869–77.
- Rabiner, L., 1989. A Tutorial on Hidden Markov Models and Selected Applications in Speed Recognition. *Proceedings of the IEEE*, 77(2), pp.257–286.
- Schmidt, T. & Wiil, U.K., Identifying Patients at Risk of Deterioration in the Joint Emergency Department. Cognition, Technology & Work. Under review.
- Sijs, H. Van Der et al., 2006. Overriding of drug safety alerts in computerized physician order entry. *Journal* of the American Medical Informatics Association, 13(2), pp.138–147.
- Sittig, D.F. & Factor, M., 1990. Physiologic trend detection and artifact rejection: a parallel implementation of a multi-state Kalman filtering algorithm. *Computer methods and programs in biomedicine*, 31(1), pp.1–10.
- Subbe, C.P. & Welch, J.R., 2013. Failure to rescue: using rapid response systems to improve care of the deteriorating patient in hospital. *Clinical Risk*, 19(1), pp.6–11.
- Sweeting, M.J., Farewell, V.T. & De Angelis, D., 2010. Multi-state Markov models for disease progression in

the presence of informative examination times: an application to hepatitis C. *Statistics in medicine*, 29(11), pp.1161–74.

- Tarassenko, L., Hann, a & Young, D., 2006. Integrated monitoring and analysis for early warning of patient deterioration. *British journal of anaesthesia*, 97(1), pp.64–8.
- Wei, H., He, J. & Tan, J., 2011. Layered hidden Markov models for real-time daily activity monitoring using body sensor networks. *Knowledge and Information Systems*, 29(2), pp.479–494.
- Windle, J. & Williams, J., 2009. Early warning scores: are they needed in emergency care? *Emergency nurse*: the journal of the RCN Accident and Emergency Nursing Association, 17(2), pp.22–6.
- Zegers, M. et al., 2009. Adverse events and potentially preventable deaths in Dutch hospitals: results of a retrospective patient record review study. *Quality & safety in health care*, 18(4), pp.297–302.
- Zeng, J., Duan, J. & Wu, C., 2010. A new distance measure for hidden Markov models. *Expert Systems* with Applications, 37(2), pp.1550–1555.
- Zhang, Y., Silvers, C.T. & Randolph, A.G., 2007. Realtime evaluation of patient monitoring algorithms for critical care at the bedside. In *Engineering in Medicine* and Biology Society, 2007. EMBS 2007. 29th Annual International Conference of the IEEE. pp. 2783–6.
- Zmiri, D., Shahar, Y. & Taieb-Maimon, M., 2012. Classification of patients by severity grades during triage in the emergency department using data mining methods. *Journal of evaluation in clinical practice*, 18(2), pp.378–88.