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A DECISION SUPPORT SYSTEM FOR THE MONITORING OF PATIENTS TREATED BY HEMODIALYSIS BASED ON A BAYESIAN NETWORK

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Abstract: Telemedicine is a mean of facilitating the distribution of human resources and professional competences. It can speed up diagnosis and therapeutic care delivery and allow peripheral healthcare providers to receive continuous assistance from specialized centers. The need of specialized human resources becomes critical with the aging of the population. The treatment of renal failure is an example where telemedicine can help to increase care quality. Over the last decades Bayesian networks has become a popular representation for encoding uncertain expert knowledge. Dynamic Bayesian networks are an extension of Bayesian networks for modeling dynamic processes. We developed a dynamic Bayesian network adapted to the monitoring of the dry weight of patients suffering from chronic renal failure treated by hemodialysis. An experimentation conducted at dialysis units indicated that the system is reliable and gets the approbation of its users.

Introduction

Telemedicine is the use of telecommunication technologies for medical diagnosis and patient care. It can deliver health care services when the provider and the client are separated by distance. Health care professionals can now access or exchange information for diagnosis, treatment and prevention of diseases and injuries [1]. Hemodialysis is the most common method used to treat advanced and permanent kidney failure. It is a heavy treatment requiring equipments and surveillance. Telemedicine can help to improve care quality for patients treated at home or at autodialysis units. The medical action depends on the doctors reasoning capacity and his aptitude to make decisions relying on potentially uncertain information.

A first approach to deal with the problem of knowledge representation in an uncertainty context is to use probability theory. Bayesian networks have become a popular representation in Artificial Intelligence for encoding uncertain knowledge [2][3].

In this article, we develop a decision support system for the surveillance of patients suffering from renal failure and treated by hemodialysis. After a brief introduction to Bayesian networks, we present the medical problem in which we are interested. Then we describe a

Bayesian network model for the monitoring of the dry weight in hemodialysis and the different stages of the medical data analysis. Finally, we evaluate the potential of our approach by presenting some experiments and results.

Materials and Methods

Bayesian networks

Bayesian networks [3] have become a popular representation in Artificial Intelligence for encoding uncertain knowledge. They provide a natural tool for dealing with two problems that occur throughout applied mathematics and engineering: uncertainty and complexity[4]. Particularly they are playing an increasingly important role in many medical applications. The Bayesian networks are able to compute efficiently probability distributions and encode easily the interaction between events.

They belong to the class of graphical models. Bayesian networks are probability graphs in which the nodes are associated to random variables and the arcs represent dependencies between the variables. The structure of the Bayesian network gives a factorization of the joint probability distribution of a set of variables.

The joint probability distribution of random variables $S = \{X_1, \dots, X_N\}$ in a Bayesian network is calculated by the multiplication of the local conditional probabilities of all the nodes. Let a node X_i in S denote the random variable X_i , and let $Pa(X_i)$ denote the parent nodes of X_i . Then, the joint probability distribution of $S = \{X_1, \dots, X_N\}$ is given as follows:

$$P(X_1, X_2, \dots, X_N) = \prod_{i=1}^N p(X_i | Pa(X_i))$$

Dynamic Bayesian networks

Dynamic Bayesian Networks (DBN) have been introduced to represent the time dimension of random variables. They are a generalization of Kalman Filter Models (KFM) and Hidden Markov Models (HMM)[5]. In the case of a (HMM), the hidden state space can be represented in a factored form instead of a single discrete variable.

A DBN encodes the joint probability distribution of a time-evolving set of variables $X[t] = \{X_1[t], \dots, X_N[t]\}$.

Usually DBNs are defined using the assumption that $X[t]$ is a first order Markov process [4]. Using the factorization property of Bayesian networks [4][6], the joint probability density of $X^T = (X[1], \dots, X[T])$ can be written as:

$$P(X[1], \dots, X[T]) = \prod_{t=1}^T \prod_{i=1}^N p(X_i[t] | Pa(X_i[t]))$$

where $Pa(X_i[t])$ denotes the parents of $X_i[t]$.

Application context

The human body gains water by drinking, eating but it also has a metabolic production of water. The water is eliminated by respiration, defecation, sweating and urination, which is the only regulated loss. The kidney is responsible for keeping constant the amount of water as needed by the organism. The progressive or sudden loss of this ability is known as kidney failure. Chronic kidney failure treatments are renal dialysis and kidney transplant.

By doing dialysis, the nephrologist intends to maintain his patient in a normally hydrated state which corresponds to a reference body weight for the patient known as the dry weight. A major difficulty of the treatment is the estimation of this ideal weight since the hydration level of the patient cannot be directly measured. The evaluation made by the physician is mainly based on the monitoring of the body weight and the blood pressure of the patient. A misadjusted dry weight has the consequence to overhydrate or dehydrate the patient.

A telemedicine experiment on peritoneal dialysis

In peritoneal dialysis (PD), a soft tube called a catheter is used to fill the abdomen of the patient with a cleansing liquid called dialysis solution. The walls of the abdominal cavity are lined with a membrane called the peritoneum, which allows waste products and extra fluid to pass from the blood into the dialysis solution. The wastes and fluid then leave the body when the dialysis solution is drained.

The most common form of PD, continuous ambulatory peritoneal dialysis (CAPD), doesn't require a machine. As the word ambulatory suggests, the patient can walk around with the dialysis solution in his abdomen. Other forms of PD require a machine called a cyclor to fill and drain the abdomen, usually at nights.

A first telemedicine experiment was conducted on CAPD patients which has led to the creation of the Diatelic company. Monitored patients send daily medical data through the Internet. These data mainly include the patient weight and his blood pressure measured in both lying and standing positions. A hidden Markov model was used to evaluate the hydration state of the patient and his dry weight adjustment. The estimation of the patient state from observations was made using the Baum-Welch algorithm [7].

In a further work we used the formalism of Bayesian networks for PD monitoring which permits a finer representation of causal relationships between the variables.

The good results obtained on PD encouraged us to use Bayesian networks for developing a decision support system adapted to hemodialysis.

Hemodialysis

Hemodialysis is the most common method used to treat advanced and permanent kidney failure. In hemodialysis the blood is allowed to flow through an artificial kidney during dialysis sessions. Contrary to the CAPD, hemodialysis is not a continuous treatment. These sessions usually take place three or four times a week. As shown on figure 1, the patient gains water between the dialysis sessions and his weight is lowered to the dry weight at the end of each session.

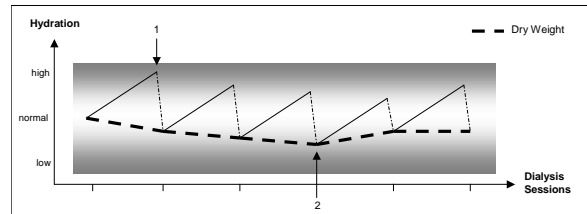


Figure 1: On this figure we see that the patient gains water between dialysis sessions and loose water during the dialysis sessions. The estimation of the dry weight is made empirically. When the patient presents signs of overhydration (1) the dry weight should be decreased. If the patient is dehydrated at the end of the dialysis session (2) the dry weight has to be increased.

DialHemo is a national INRIA action for developing a tele-surveillance and tele-diagnosis system adapted to kidney failure patients treated by hemodialysis either at home or at autodialysis units. The system will be accessible by nephrologists as well as general practitioners. Several dialysis units are now connected to the system.

The monitoring of the dry weight adjustment in hemodialysis is a regulation problem. The patient provides information on his hydration state (weight, blood pressure,...) and, by his actions (increase/decrease the dry weight), the physician modifies the hydration state of his patient. The decision support system can be considered as a third agent interacting with both the patient and the physician (see figure 2). It improves the communication of information from the patient to the physician by processing the data and selecting the information of interest. The decision support system does not replace the physician visits. It comes in addition, providing information between the visits. The processing and selection of information in a medical context is certainly a delicate matter. The information provided should be relevant. The system must not miss real alerts and, on the other hand, it should not produce too much false alerts. The purpose of our application is to monitor the dry weight of patients with chronic renal failure treated by hemodialysis, and to help the physician in his decision. It cannot replace the physician's expertise for making the decision. The responsibil-

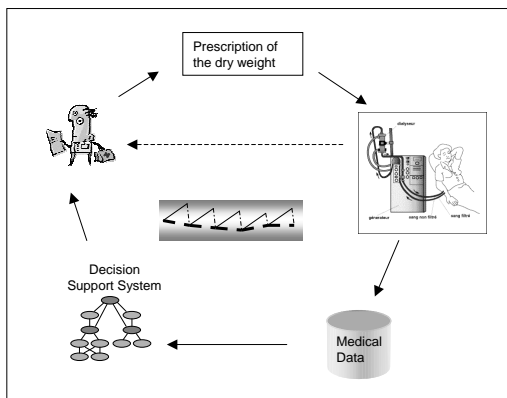


Figure 2: Information flows between the patient, the physician and the decision support system.

ity of the treatment remains in the hands of the physician.

Indicators for dry weight diagnosis

The first step in developing a decision support system is to define a sufficient set of indicators for making the diagnosis. This work was done with the collaboration of the nephrologists. We limited the data used for the diagnosis to elements which can easily be acquired without the presence of a physician since the system should be adapted to patients treated at autodialysis units. This excludes clinical signs such as edema.

Each indicator will be associated to an observed variable in the Bayesian network.

- **Blood pressure:** The more the blood flow contains water the bigger its volume is and as a consequence the force of the blood pushing against the walls of the arteries grows. High/low blood pressure can be a symptom of overhydration/dehydration. Blood pressure is measured at the beginning (BPb) and at the end (BPa) of the dialysis. Orthostatic hypotension means low blood pressure on upright posture and it can be a symptom of dehydration. The difference between blood pressure on lying and upright posture is measured before (OHb) and after (OHa) the dialysis.
- **Weight:** Except in the case where one is starving or infected, real tissue weight is not gained or lost rapidly. If the patient's weight grows by one kilogram in one week, there is a risk that it is one kilogram of water. The evolution of the patient weight is a key element for dry weight diagnosis. The weight of the patient is measured before (Wb) and after (Wa) the dialysis. At the end of the dialysis the weight of the patient should be nearly equal to the dry weight prescribed by the physician.
- **Ultrafiltration:** This quantity (UFT) is calculated as the difference between the weight at the beginning of the dialysis and the dry weight estimated by the physician. It represents the amount of water to be extracted during the dialysis. The extraction speed of the water depending on the duration of the dialysis

session is known as the ultrafiltration rate (UFR). If the (UF) / (UFR) is high the dialysis session may not be well tolerated by the patient which could result in low blood pressure at the end of the session.

In addition to the indicators we need to define variables for representing the state of the patient (which is not directly observable). The estimation of the hydration and the dry weight adjustment of the patient is represented by the following hidden variables:

- **Dry Weight** adjustment: This variable (DW) indicates if the dry weight is correctly adjusted for the patient or if it should be decreased/increased.
- **Hydration before** the dialysis session (Hb)
- **Hydration after** the dialysis session (Ha)

Table 1 summarizes the variables used in the Bayesian network we described before.

Table 1: Variables of our Bayesian network

Variables	
Hidden	Observed (indicators)
Dry Weight (DW)	Weight before (Wb)
Hydration before (Hb)	Weight after (Wa)
Hydration after (Ha)	Blood Pressure before (BPb)
	Blood Pressure after (BPa)
	Orthostatic H. before (OHb)
	Orthostatic H. after (OHa)
	Ultrafiltration (UFT)
	Ultrafiltration Rate (UFR)

A dynamic Bayesian network model for hemodialysis monitoring

The theory of Bayesian networks allows us to represent relationships between these observed and hidden variables in a probabilistic way which is well adapted to the uncertainty inherent to medical questions. Causal links between the variables are represented on figure 3.

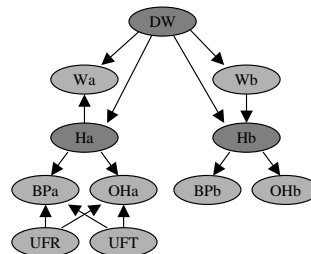


Figure 3: Static Bayesian Network for dry weight adjustment diagnosis

This Bayesian network shows the causal relations between hidden and observed variables. For example we can see that the hydration before dialysis is a cause for

both the weight before dialysis and the blood pressure before dialysis.

Observations made during a single dialysis session are not sufficient to properly evaluate the dry weight adjustment. Experiments conducted using this simple layer Bayesian network showed that the diagnosis was highly instable and thus producing false alerts. In order to represent the influence of past events over the present state of the patient, it is necessary to extend this model into a dynamic Bayesian network. We consider the dry weight as a first order Markov process so as to integrate the time dimension of this variable. The physician's action on the dry weight (increase / decrease) should also be taken into account when considering the time dimension of the variable. The diagnosis of dry weight adjustment should be based on the present dialysis session observations, the previously made estimation of the dry weight adjustment, and the potential actions made by the physician (dot lined node on figure 4).

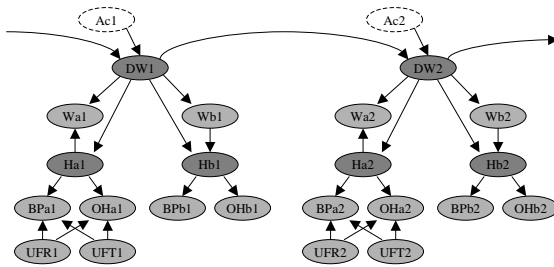


Figure 4: A Dynamic Bayesian Network model for dry weight adjustment diagnosis

In our Bayesian network, nodes representing the hydration state of the patient (Hb) and (Ha) are not connected between successive layers (corresponding to successive dialysis sessions). This choice can be justified as follows: contrary to the dry weight, the hydration is highly sensible to daily variations depending for example on the patients meals, the temperature or the physical efforts made by the patient. The influence of these external causes makes a causal link between the hydration variables of successive sessions irrelevant.

Our Bayesian network is based on discrete variables. Its structure as well as its parameters are based on human expertise. They have been defined with the collaboration of nephrologists. The dynamic Bayesian network data input is pre-processed in order to convert the medical information into a probability distribution over the network variables. The observations made over the three past sessions are entered as evidences and propagated into the dynamic Bayesian network with the exact inference algorithm JLO [8]. Thus we get the *a posteriori* probability distributions over the hidden variables of the network.

A medical data acquisition platform

A software was designed by the Diatic company to collect medical data from dialysis units participating to

the experiment. For units previously using a data management software, the experimental platform was interfaced with it. For units not using a data management software, a new one was designed. The Diatic platform has the following features:

- Display of patients data
- Display of decision support system output and automated email alerts generation
- Acquisition and display of medical prescriptions
- Acquisition and display of biological analysis
- Integrated message system

Preprocessing of medical data

In order to be entered as soft-evidences into the dynamic Bayesian network, daily measures are transformed with fuzzy sigmoid operators into likelihoods to be greater than, equal to or lower than a given base. This preprocessing can be decomposed into three stages:

- **Centering:** The first stage consists in centering the measure around a reference value considered as normal for the patient. This reference is calculated based on statistics on the value's record history. For example the blood pressure is centered around the average blood pressure calculated for the patient over the last 15 days. The normal value for the weight at the end of the dialysis session (W_a) is the dry weight. This stage can be considered as an adaptation of the model input to the patient's data.
- **Normalization:** A tolerance is defined for each medical measure (weight, blood pressure, ultrafiltration) in order to normalize the values between -1 and 1 . This tolerance defines the width of the measure interval corresponding to the normal state of the discrete variable. When the value is saturated (equal to 1 or -1), it is a certainty that the variable is in state lower than or greater than the base.
- **Discretization:** The discretization consists in converting the normalized value into likelihoods. In our case, the variables of the Bayesian network have three states: *low*, *normal*, and *high*. The corresponding likelihoods are given by three fuzzy filters (see representation figure 5). The use of fuzzy filters has the advantage of preventing threshold effects.

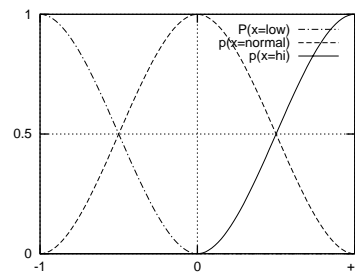


Figure 5: Probability density functions

Interpretation of a posteriori probabilities

The diagnosis produced by the Bayesian network is the *a posteriori* probability distribution of the hidden variable DW . The belief that the dry weight should be increased/decreased is given by the probability distribution $P(DW|observations)$.

We used a simple threshold alarm for generating alerts. The system generates an alert when the probability $P(DW = high)$ or $P(DW = low)$ is greater than 0.5 for two consecutive sessions. This value corresponds to the point where $P(DW = high/low)$ becomes greater than $P(DW = normal)$. Figure 6 shows the influence of the alert threshold on the number of generated alerts.

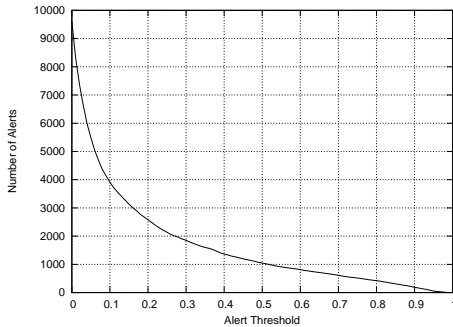


Figure 6: Influence of the threshold (th) on the number of generated alerts for 9600 recorded dialysis sessions. For $th < 0.5$ the number of alerts is decreasing exponentially and for $th > 0.5$ the decrease becomes linear.

The physician has the possibility to consult the system diagnosis which is presented as a single curve for simplification reasons. This curve is obtained by subtracting $P(DW = low)$ from $P(DW = high)$. This gives a representation between -1 and 1 indicating that the dry weight is low, normal or high. See figure 7 for an example of diagnostic curve.

Results

An experimentation has been conducted for more than a year in dialysis units. The dialysis sessions of about one hundred patients have been monitored by our decision support system.

Figure 7 shows an example of a diagnosis output over three months for a patient treated with hemodialysis. It shows the diagnosis and alerts compared to the physician's decisions. Each dot corresponds to an email sent to the physician alerting him of a high/low dry weight. Arrows correspond to sessions where the physician decided to increase or decrease the dry weight. Between 28/11/2004 and 23/01/2005 several alerts were generated indicating that the dry weight was too high. These alerts were approved by the physician who indeed decreased the dry weight several times. On 25/02/2005 a low dry weight alert was generated followed by an increase of the dry weight by the physician.

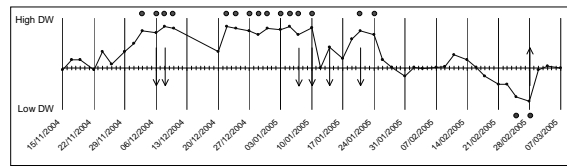


Figure 7: Monitoring of the dry weight adjustment of a patient treated by hemodialysis

Discussions

Conclusions of nephrologists participating at the experimentation are the following. Primary results on care quality evolution are encouraging. The program gets the approbation of users not directly involved in the experimentation. Patients are satisfied to have access to their session data, they feel under better surveillance and are more respectful of medical prescriptions. The expert system is reliable and in agreement with medical decisions most of the time. It seems that some signals, directly given by the dialysis device (like the *hemoscan* which measures blood volume variations during the dialysis session), could help the expert system to anticipate the evolution of the dry weight.

Conclusions

In this paper, we described an application of dynamic Bayesian networks to the monitoring of patients treated by hemodialysis. The purpose of this decision support system is to help the physicians with the estimation of the dry weight.

The diagnosis as we presented it in this article is obtained from the patient data (weight, blood pressure...) as well as the data of the dialysis session (dry weight, duration...). A pre-treatment is used in order to transform medical data into standardized data usable by the system.

The analysis of care quality gives good results. Alerts generated by the expert system are reliable and its diagnosis is most of the time in agreement with the physician's decisions. Signals directly given by the dialysis device will be added to the expert system. It seems that some of these signals could help to anticipate the evolution of the dry weight.

In spite of the good results of our system, the estimation of the dry weight remains a rather complex problem even with the physicians' experience. The value of the dry weight is subjected to variations depending on the evolution of the patient's body fat mass. The indicators which permit the diagnosis of the dry weight (as for example blood pressure) are subjected to various external factors (meals, drugs...). All these external factors make the estimation of the dry weight very difficult.

Future work relates to the following points:

- The parameters used are generic, and therefore badly adapted to the particular cases, as for example heart failure patients. The generic model could be adapted

to each patient by the definition of personalized profiles. These specific models would permit a finer diagnosis of the dry weight adjustment.

- The dialysis machine has several sensors which provide data throughout the dialysis session. The next development stage is to integrate some of them in the system analysis. We will focus on the data which gives information about the water volume in the organism like the *hemoscan*. The integration of such elements could help to anticipate the evolution of the dry weight.

Acknowledgment

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