Sharing and Mining of Rare Drug Reactions and Symptoms

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ABSTRACT—Adverse Drug Reaction (ADR) is one of the greatest consequence in the evaluation of drug safety. Also, most of the adverse drug reactions are not discovered during limited pre-marketing clinical trials; but, they are only observed only after a long term post-marketing observation of drug usage. The exposure of adverse drug reaction, is an important method of research technique for the pharmaceutical industry. Recently, more number of adverse events and the improvement of data mining technology have motivated the development of statistical and data mining methods for the detection of Adverse drug reactions. These methods, without integration into the knowledge discovery systems, are very tedious and uncomfortable for users and the processes for exploration are time-consuming.

Index Terms—Adverse drug reactions, data mining methods.

1. INTRODUCTION

To find causal associations between two sets of events which has low frequency is very useful for numerous real-world applications. Even if the probability is low, a drug that is used at suitable dose may cause more than one adverse drug reactions (ADRs). This type of causal relationships helps to correct negative results caused by its antecedents. In this paper, we try to make use of a knowledge-based approach to find the degree of causality of an event pair. This happens within the sequence since the intention of causality is frequently application dependent. Even though the new framework can be applied to many implementations, our study was stimulated through the need of finding ADR signals in postmarketing observation. Detecting Signals of Adverse Drug Reactions from Health Consumer Contributed Content in Social Media [1].

ADRs is a serious world-wide problem and can intricate a patient’s medical condition even death. It contribute about 5 percentage of every hospital admissions and has the fifth usual cause of death in hospitals. Clinical trials are required for premarketing, these trails are not capable of finding rare ADRs. Postmarketing has drug safety that is monitoring of medicines once they have been available to customers. ADRs reports are filled at the judgment of the users which has gross under reporting [3], [7]. Data mining algorithm which is known as Multi-item Gamma Poisson Shrinker [5] is used for finding potential signals from its spontaneous reports.
An electronic patient report[2] become more easily accessible in different health organizations to provide a new source of information that can generate ADRs signals much earlier[6]. An important measure known as the causal-leverage was created on the method of calculating fuzzy RPD[3].

2. RELATED WORKS

Many measures have been proposed to mine association rules in the form of X and Y. Support and confident are the original measures proposed for association rule. Causal modeling have been widely studied in the field of statistics theory[8]. The eventsets rules support lesser than threshold and it is implemented in different strategies[9].

This paper suggests an interactive system proposal for the exposure of ADRs. By combining an ADR data warehouse and original data mining procedure, the proposed system not only supports OLAP style multidimensional study of ADRs, however allows the interactive invention of associations between drugs and symptoms, known as a drug-ADR relationship rule, which can be additional developed using some other factors of attention to the user, like demographic information.

The research shows that remarkable and important drug-ADR association rules can be resourcefully mined. Finding the connection between two event sets is an important area in data mining research. One of the most famous models is Pearl’s causal model where causal modeling are based on representations by directed acyclic graphs (DAGs). For a more detailed review about the state of the art of causality readers are referred to a recent survey paper [10]. With a high rise of interest in association rule mining in the latest years, some of the exceeding model have been borrowed to extract the causal relationships between the two events or event sets.

2.1 Detecting Signals of Adverse Drug Reactions

In this paper [1], Adverse drug reactions are causing a substantial amount of hospital admissions and deaths, which cannot be underestimated. Although a great effort has been put on the pre-marketing review during pharmaceutical product development, it cannot identify all possible adverse drug reactions. Currently, post-marketing surveillance is conducted through centralized volunteering reporting systems.

2.2 Temporal pattern discovery in longitudinal electronic patient records

Large collections of electronic patient records provide a vast but still underutilized source of information on the real world use of medicines [2]. They are maintained primarily for the purpose of patient organization, but contain a wide range of clinical in sequence highly related for data analysis. While they are a normal resource for epidemiological positive studies, their use in context of gropping data analysis is still incomplete. Here, we present a structure for open-ended pattern detection in large patient records repositories.

2.3 Proactive Post marketing Drug Safety Surveillance

Discovering unknown adverse drug reactions (ADRs) in postmarketing surveillance as early as possible is of great importance [3]. The present approach to postmarketing surveillance mainly relies on unstructured reporting. It is an inactive surveillance system and partial by gross underreporting (<10% of reporting rate), latency, and conflicting reporting. We propose a novel team-based intellectual agent software system
approach for proactively monitoring and detecting potential ADRs of interest using electronic patient records.

Pattern mining techniques are designed for extracting interesting and useful item sets[4] from a database. Amongst them, frequent item sets are usually thought to unfold “regularities” in the data, in which they are the witnesses of regular phenomena and they are consistent along with the correct expectation of the domain ex. In some situations however, it may be interesting to search for “rare” items etc, i.e. item sets that do not occur frequently in the data (contrasting frequent item sets). These correspond to unpredicted phenomena, probably contradicting beliefs in the domain expert.

2.5 Conditions on Fuzzy RPD Model

RPD model gives a famous cognitive assessment model which is useful for showing decisions of human experts based on their previous experiences[5]. Experience plays an important role in RPD model and they are stored in experience knowledge base (EKB).

3. SEARCHING FOR ADR SIGNAL PAIRS

We developed and incorporated an exclusion mechanism that can effectively reduce the unwanted effects caused by regular events. Here, our new measure is called exclusive causal-leverage measure. We future a data mining algorithm which is to mine the ADR signal pairs from the electronic patient database based on its new measure. The algorithm’s computational density is to be analyzed. We compared our latest exclusive causal-leverage measure with our earlier proposed causal-leverage measure and also two traditional measures in the literature: leverage and risk ratio.

Searching for the drugs and support count for the each drug

- drugHashTable ¼ null
- for each patient Pk 2 DB do
  - retrieve all the drugs Dk taken by the patient
  - for each drug dkl 2 Dk do
    - if (drugHashTable:containsKey(dkl) == false) do
      - Sigma= 1 {a new drug dkl is found and set its Support count as 1}
    - else
      - Sigma=drugHashTable:getValue(dkl) + 1 {update Support count}
    - end if
- drugHashTable.putValue(dkl,Sigma)
- end for
- end for
- return (drugHashTable)

To create the superiority of our latest measure, we did wide-ranging experiments. In our earlier work, we experienced the effectiveness of the causal-leverage calculate using a single drug in the research. In this
paper, we chosen three drugs and evaluate the top 10 ICD-9 (International Classification of Diseases, ninth Revision) codes ranked by exclusive causal-leverage measure for the each drug. We also test how the length of hazard phase affects the presentation of the exclusive causal-leverage measure. Support count for each drug or the symptom will be used to calculate the exclusive causal-leverage value for its related pairs. We can find the power of the causality between a drug and a exacting possible ADR. Data mining algorithm called Multi-item Gamma Poisson Shrinker.

4. ALGORITHM

4.1 Searching for the drugs and the support count for each drug.

This algorithm explains the way to explore a database for the drug lists and the support count for the each drug. The exposed drugs and theirs sustains are saved in the hash table. The support count for each drug or symptom will be used to calculate the exclusive causal-leverage value for related pairs. We can find the Strength of the causality between a drug and a particular possible ADR.

4.2 Pair Generation and Evaluation

This algorithm is used to find the Patient Symptom table, even users can also find the list of symptoms.

4.3 Procedure Causal-leverage(X,Y,PID)

It Shows the way to compute Causal-leverage value of a general pair between X and Y.

5. CONCLUSION

Mining the causal association between two events is very important and useful in many real applications. It can also help people to discover the causality of a type of events and avoid its potential adverse property. Though, mining these associations is very hard, especially when events of interest occur occasionally. We have developed a new interestingness measure, exclusive causal-leverage, based on the knowledge-based fuzzy RPD model. This method can be used to quantify the degree of association of a CAR. Also, the measure was designed to mask the undesirable effects caused by high regularity events. We have useful this measure to detect the causal associations between each of the three drugs (i.e., pravastatin, rosuvastatin, and enalapril,) and the ICD-9 codes. A data mining algorithm was developed to search a real electronic patient database for possible ADR signals. New results showed that our algorithm could effectively make known ADRs rank high among all the symptoms in the database.

6. REFERENCES


