Weighted Markov chains for forecasting and analysis in Incidence of infectious diseases in Jiangsu Province, China☆

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Abstract

This paper first applies the sequential cluster method to set up the classification standard of infectious disease incidence state based on the fact that there are many uncertainty characteristics in the incidence course. Then the paper presents a weighted Markov chain, a method which is used to predict the future incidence state. This method assumes the standardized self-coefficients as weights based on the special characteristics of infectious disease incidence being a dependent stochastic variable. It also analyzes the characteristics of infectious diseases incidence via the Markov chain Monte Carlo method to make the long-term benefit of decision optimal. Our method is successfully validated using existing incidents data of infectious diseases in Jiangsu Province. In summation, this paper proposes ways to improve the accuracy of the weighted Markov chain, specifically in the field of infection epidemiology.

Key words: weighted Markov chains, sequential cluster, infectious diseases, forecasting and analysis, Markov chain Monte Carlo

INTRODUCTION

Mathematical models of any natural phenomenon should rest on some basic knowledge of the phenomenon and the data collected to track and understand it. Many years ago, J.L. Doob had defined a "stochastic process" as the mathematical abstraction of an empirical process whose development is governed by probabilistic laws. It is important to note that the term "stochastic process" refers to the mathematical abstraction, model, or representation of the empirical process and not to the empirical process itself. During recent years, the theory of stochastic process has developed very rapidly and has found application in a large number of fields.[1]

In particular, a class of stochastic processes termed Markov chains or processes has been investigated extensively. Markov chains are one of the richest sources of models for capturing dynamic behavior with a large stochastic component.[2,3] It is of great importance in many branches of science and engineering and in other fields, including physics[4,5].
工业控制\[^{[6,7]}\]，可靠性分析\[^{[8]}\]，优化分析\[^{[9]}\]，经济\[^{[10,11]}\]，等。马尔科夫链理论方法是定量分析在情况中转移系统的关键，从一个状态到另一个状态，因此预测未来趋势。这为做出战略决策提供了基础。

在医学和公共卫生领域，疾病的发生、发展和预后将不可避免地受到外部因素和人体因素的影响。这些因素是紧密相关的，一个因素的变动将影响另一因素，这很难用结构化的原因-结果模型来解释。然而，以变量之间互为依存关系来描述这些数据，是最重要的和有用的特征，为研究目标提供了有效的途径。这里，我们将建立一种动态模型来描述这种依存关系。

在过去，许多学者应用马尔科夫链理论来预测传染病的发生，建立了一些相应的数学模型。这种方法可以用于分析传染病的长期趋势、季节特征、周期性、短期波动和不规则变化。这些特征包括长期趋势、季节特征、周期性、短期波动和不规则变化。

在该领域，一些研究已用于分析和分类，如在江苏省的研究，以及在更广泛的领域中应用。这些方法可以根据历史数据，具有较高的准确性，因为它们依赖于外部因素，具有广泛的应用适应性。然而，这些方法的复杂性和准确性，使得它们在实际应用中具有挑战性。因此，需要进一步研究和改进这些方法，以提高其准确性和可靠性。

**ONE-DIMENSIONAL SEQUENTIAL CLUSTER ANALYSIS**

聚类分析涉及产生聚类的技术，这些技术生产聚类，而不是初始未分类的数据。这种分类可能是基于数据中固有的特征，而不是被指定的特征。基于疾病的发生率数据的特征，本文试图使用一维顺序聚类分析算法来衡量数据的聚类效果。

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the diameter of that section, denoted as \( D(i, j) \):

\[
D(i, j) = \sum_{l=1}^{j} (x_l - x_j)(x_l - x_0) \quad (1.2)
\]

Divide \( n \) sequential variants into \( k \) kinds, and any partition can be

\[
P(n,k) = \{i_1,i_1+1,\ldots,i_2-1,i_2,i_2+1,\ldots,i_3-1,\ldots,i_k \}
\]

Define the error function, namely the objective function of this partition, and let it be the total sum of squares of deviations in this kind:

\[
L[P(n,k)] = \sum_{j=1}^{k} D(j,i,j+1) - 1 \quad (1.3)
\]

When \( n \) and \( k \) are fixed, the smaller the error function \( L[P(n,k)] \) is, the smaller the sum of squares of deviations within each kind, and this proves the reasonability of the classification. It can be proved that the so-called optimum partition is to make the \( L[P(n,k)] \) smallest. \( k \) can be calculated according to the relation curve of \( L[P(n,k)] \) and \( k \). The value at the turn of the curve is the optimum partition number.

**WEIGHTED MARKOV CHAIN**

A stochastic process \( X = \{X(t), t \in T \} \) is a collection of random variables. That is, for each \( t \) in the index set \( T \), \( X(t) \) is a random variable. We often interpret \( t \) as time and call \( X(t) \) the state of the process at time \( t \). If the index set \( T \) is a countable set, we call \( X(t) \) a discrete-time stochastic process, and if \( T \) is a continuum, we call it a continuous-time stochastic process. The collection of possible values of \( X(t) \) is called state space. This general model has been described, from a theoretical analysis, by Chiang [3] and others [3].

**Markov chain**

Markov chain is a branch of Markov process. If the present state of the system is given, then the past and future are (conditionally) independent. Such a behavior is called the Markov property of the system. A Markov chain evolves in a discrete (countable) state space with respect to discrete or continuous time.

A stochastic process \( X = \{X(t), t \in T \} \) is defined on a probability space \((\Omega, F, P)\), where parameters set \( T = \{0,1,2,\ldots\} \), and state space \( E = \{0,1,2,\ldots\} \). It is called a Markov chain if for any positive integers \( l,m,k \) and \( j_1 < \ldots < j_k \), \( \mu_{l,m,k} = \mu_{l,k} \mu_{m,k} \mu_{l,m} \mu_{k,l} \in E \),

\[
P[X(m+k) = i_{m+k} | X(m) = i_m, X(j_1) = j_1, \ldots, X(j_k) = j_k] = P[X(m+k) = i_{m+k} | X(m) = i_m]
\]

For the aperiodic Markov chain, we have

\[
\pi_j = \frac{1}{\mu_{ij}} \quad (2.2)
\]

where \( \mu_{ij} \) denote the mean recurrence time to state \( j \), and \( \pi_j \) is the limiting probability. The preceding identity shows that one way to find the limiting probability is by taking the reciprocal of the mean recurrence time. A simple way to find \( \{\pi_j\} \) will be given shortly.

When an irreducible Markov chain is aperiodic and positive recurrent, the chain is called an ergodic Markov chain. The limiting distribution \( \{\pi_j\} \) of an ergodic chain is the unique nonnegative solution of Equations:

\[
\pi_j = \sum_{k=0}^{\infty} \pi_k P_{kj} \quad j = 0, 1, 2, \ldots (2.3)
\]

Now \( \pi_j \) may be interpreted as the long-run proportion of time that the Markov chain is in state \( j \). Thus it is easily seen to satisfy (2.2). The solution of these equations, sometimes, is not straightforward, and the MCMC methods may be used to solve them [32], which is considered in the next section.

There are many properties and relative conclusions about Markov chain, and some other mathematical expressions (e.g., recurrent, limit theorems, periodic, etc.) are described by Freedman [33] and Kendall and Montana [34].

**Weighted Markov chain**

Because the monthly (or yearly, weekly) incidence of infectious disease are a series of correlative random variables, self-correlation coefficients depict various disease incidence data relationships. The past several months’ incidence of infectious disease can be considered in advance to predict the present month incidence data. Then the weighted average can be made according to the incidence of the past several months infectious diseases compared with the present month’s. Therefore the prediction purpose to make full and rational use of information is reached. That is the basic thought of weighted Markov chain prediction.

Based on the above discussion in this paper, the specific method of weighted Markov chain prediction is expressed as follows:

1. Set up a classification standard of the monthly incidence of infectious disease according to the length of material series and the requirement of the specific problems. For instance, we can classify incidence of infectious disease as one-dimensional sequential cluster analysis in section 2 (corresponding to state space \( E = \{1, 2, 3, 4, 5, 6\} \)) and so on.
2. Determine every month’s incidence of infectious
disease state according to the classification standard of 
“①”.
③ Compute various self-correlation coefficients $r_k$, $k \in F$
\[ r_k = \frac{\sum_{i=1}^{n-k} (x_i - \bar{x})(x_{i+k} - \bar{x})}{\sum_{i=1}^{n} (x_i - \bar{x})^2} \quad (2.4) \]
where $r_k$ indicates $k$ months self-correlation coefficient, $x_i$ ($i=1,2,\cdots,n$) indicates the $i$th months infectious disease incidence, $\bar{x}$ indicates the mean value of $x_i$, $n$ indicates the length of monthly incidence of the infectious disease series.
④ Standardize various self-correlation coefficients. In other words, that is take
\[ w_k = \frac{|r_k|}{\sum_{k=1}^{m} |r_k|} \quad (2.5) \]
as weights of various (steps) Markov chain ($m$ is the maximum step according to prediction).
⑤ According to statistical results of “③”, we can get various steps of Markov chain transition probabilities matrixes, which decided the probability law when incidence of infectious disease states transited.
⑥ The past several months incidence of infectious disease can be initial states respectively, the state probability of the present month’s incidence of infectious disease $P_i^{(k)}$, $i \in E$ can be predicated and combined with relative transition probabilities matrixes, $k$ indicates the step of Markov chain, $k=1,2,\cdots,m$.
⑦ Take the weighted average of various predicting probabilities of the same state as predicting probability of the plum rains intensity index, that is
\[ P_i = \sum_{k=1}^{m} w_k P_i^{(k)}, \quad i \in E. \quad (2.6) \]
If $P_i = \max\{P_i, \ i \in E\}$, $i$ is the predicting state of the present month’s incidence of infectious disease. After the present month’s incidence of infectious disease is determined, we can add it to the original series, repeating steps "①-⑦", and the next month’s incidence of infectious disease can be predicted.
⑧ The further analysis of Markov chain’s characteristics (ergodic property, stationary distribution, etc.) also can be carried out.

**MCMC METHODS**

In this section we will describe MCMC methods for the weighted Markov chains. Our approach is analogous to the one used for solving the equations (2.3) in the previous section. Since there has been extensive research conducted and written about MCMC methods, we will be brief [37]. However, it should be noted that the full posterior distribution over all parameters in the model is unwieldy.

One standard method for constructing a Markov chain with the correct limiting distribution is via a recursive simulation of the so-called full conditional densities: that is, the density of a set or block of parameters. Each of the full conditional densities in the simulation is then sampled either directly (if the full conditional density belongs to a known family of distributions) or by utilizing a technique such as the Metropolis-Hastings (M-H) method. An important and crucial point is that these methods do not require knowledge of the intractable normalizing constant of the posterior distribution.

In the present case, we applied MCMC methods to solve the above equations (2.3), iterative and computational details are described in the recent papers of Chib and Winkelmann [39] and Covington et al. [39].

**APPLICATION**

In order to explain specific applications of this method and to conduct testing, this research is based on the samples of the monthly surveillance data of Hepatitis B patients in the period of January 1980 to October 2006 in Jiangsu Province. The weighted Markov chain theory was used to make a forecast and other related analysis of the incidents of the disease in November and February 2000.

Liver cancer is one of the most life-threatening cancers, and is the third-leading cause of death from cancer in China, and the top leading cause in the Province of Jiangsu. There are some 260,000 new cases of liver cancer each year throughout the world. Of all these cancer sufferers, about 42.5% are from China, and 90% of all liver cancer patients have previously been infected by Hepatitis B virus (HBV). A collection of data we gathered and analyzed suggests that about 25% of all those infected with HBV will eventually die of chronic severe hepatitis, cirrhosis of liver and liver cancer. Moreover, both acute and chronic Hepatitis B patients are the main source of infection for HBV. China is densely populated with Hepatitis B patients. According to a nationwide hepatitis epidemiological survey conducted in 2004, the average HBV infection rate of China is 70%-90% (including people infected and being infected). Therefore, the forecasting research of the incidence of HBV has far-reaching implications.

Our forecasting and analysis study is as follows:
① Set up a classification standard of the monthly incidence of infectious disease according to the one-dimensional sequential cluster analysis algorithm by SAS 9.1.3 software. The value at the turn of the curve is $k = 4$ (see, e.g., Fig. 1).
② As Table 1 shows, the incidence data of
infectious disease can be classified into 6 grades (corresponding to 4 states of weighted Markov chain), so various months’ incidence of infectious disease states can be determined.

③ According to the Table 1 classification standard, various self-correlation coefficients and Markov chain weights of various steps can be computed (Table 2).

④ After statistical computation, various one-step transition probabilities matrices with step’s length 1, 2, 3, 4, 5 and 6 respectively were constructed:

⑤ We took the infectious disease incidence of July 1999 - Dec 1999’s series to predict the Jan 2000’s series to predict the Jan 2000’s infectious disease incidence state. The results are shown below in Table 3.

### Table 1 Classification of incidence of infectious disease for Jiangsu Province

| State | Incidence interval |
|-------|--------------------|
| 1     | \( x \leq 1029 \)  |
| 2     | \( 1029 < x \leq 1369 \) |
| 3     | \( 1369 < x \leq 1641 \) |
| 4     | \( 1641 < x \leq 1777 \) |
| 5     | \( 1777 < x \leq 2071 \) |
| 6     | \( x > 2071 \)    |

### Table 2 The weights of various steps Markov chain and various self-correlation coefficients

| \( k \) | 1 | 2 | 3 | 4 | 5 | 6 |
|---------|---|---|---|---|---|---|
| \( r_k \) | 0.4145 | 0.36038 | 0.1122 | -0.0895 | -0.09406 | -0.09895 |
| \( w_k \) | 0.3570 | 0.3104 | 0.0967 | 0.0697 | 0.0810 | 0.08552 |

### Table 3

| \( \mathbf{P}(n, k) \) | \( \mathbf{P}(n, 1) \) | \( \mathbf{P}(n, 2) \) | \( \mathbf{P}(n, 3) \) | \( \mathbf{P}(n, 4) \) | \( \mathbf{P}(n, 5) \) | \( \mathbf{P}(n, 6) \) |
|------------------------|------------------------|------------------------|------------------------|------------------------|------------------------|------------------------|
| \( \mathbf{P}(n, 1) \) | \( 5/7 \) | \( 1/14 \) | \( 1/7 \) | \( 2/7 \) | \( 1/14 \) | \( 0 \) |
| \( \mathbf{P}(n, 2) \) | \( 3/7 \) | \( 1/14 \) | \( 1/7 \) | \( 2/7 \) | \( 1/14 \) | \( 0 \) |
| \( \mathbf{P}(n, 3) \) | \( 1/30 \) | \( 1/13 \) | \( 1/3 \) | \( 1/10 \) | \( 1/5 \) | \( 0 \) |
| \( \mathbf{P}(n, 4) \) | \( 0 \) | \( 0 \) | \( 8/23 \) | \( 6/23 \) | \( 7/23 \) | \( 2/23 \) |
| \( \mathbf{P}(n, 5) \) | \( 4/35 \) | \( 6/35 \) | \( 8/35 \) | \( 3/7 \) | \( 2/35 \) | \( 0 \) |
| \( \mathbf{P}(n, 6) \) | \( 0 \) | \( 0 \) | \( 0 \) | \( 1/9 \) | \( 1/9 \) | \( 5/9 \) | \( 1/9 \) |
As Table 3 shows, max \( P_i, i \in E \) = 0.3734, then \( i = 3 \), and the infectious disease incidence state of Jan 2000 is 3. Corresponding infectious disease incidence data \( x \) satisfies: \( 1369 < x \leq 1641 \). The actual infectious disease incidence state of Jan 2000 in Jiangsu Province is 1390, and the intensity state is 3. The prediction is correct.

Similarly, the Aug 1999 - Jan 2000 month series can be used to predict the infectious disease incidence state for Feb 2000. This forecasting process is just a repeat of "①-⑤". The prediction results are listed below in Table 4.

Further analysis of this weighted Markov chain’s characteristics can be carried out as in Table 5.

From Table 5, we may infer that the return period of the state \( j \) is \( T_j \). The return period of each state will be \( T_1 = 17.14(\text{months}) \), \( T_2 = 7.5(\text{months}) \), \( T_3 = 4.14(\text{months}) \), \( T_4 = 5(\text{months}) \), \( T_5 = 3.43(\text{months}) \), and \( T_6 = 13.33(\text{months}) \) respectively. Thus it can be seen that, according to the classifying criteria determined in this article, the state of the number of incidents of Hepatitis B is most probable to appear about 3.43 months per time on average, and at 0.2917 percentage rate. The state 3 is the second, about 4.14 months per time on average, and the percentage is about 0.2417. States 4 and 2 are much less probable than the above; and the state 6 and 1 are least probable to appear, about 13.33 and 17.14 months respectively, with percentages of 0.0750 and 0.0583, respectively.

### Table 3 Infectious disease incidence state prediction in Jan 2000

| Initial year | State | Step (month) | State Weights | 1 | 2 | 3 | 4 | 5 | 6 | Probability source |
|--------------|-------|--------------|---------------|---|---|---|---|---|---|-------------------|
| Dec 1999     | 1     | 1            | 0.3570        | 0 | 0 | 5/8 | 0 | 3/8 | 0 | \( P^{(1)} \) |
| Nov 1999     | 2     | 2            | 0.3104        | 1/14 | 1/14 | 2/7 | 1/7 | 5/14 | 1/14 | \( P^{(2)} \) |
| Oct 1999     | 2     | 3            | 0.0967        | 0 | 4/13 | 0 | 2/13 | 5/13 | 2/13 | \( P^{(3)} \) |
| Sept 1999    | 3     | 4            | 0.0697        | 0 | 7/29 | 3/29 | 3/29 | 10/29 | 1/29 | \( P^{(4)} \) |
| Aug 1999     | 4     | 5            | 0.0810        | 3/22 | 3/11 | 3/11 | 0 | 7/22 | 0 | \( P^{(5)} \) |
| Jul 1999     | 4     | 6            | 0.0852        | 4/21 | 4/21 | 5/21 | 2/21 | 5/21 | 1/21 | \( P^{(6)} \) |
|              |       |              | \( P_i \) (weighted average) | 0.0495 | 0.1071 | 0.3734 | 0.0745 | 0.3520 | 0.0435 |

### Table 4 Infectious disease incidence state prediction in Feb 2000

| Initial year | State | Step (month) | State Weights | 1 | 2 | 3 | 4 | 5 | 6 | Probability source |
|--------------|-------|--------------|---------------|---|---|---|---|---|---|-------------------|
| Jan 2000     | 3     | 1            | 0.3570        | 1/30 | 1/3 | 1/3 | 1/10 | 1/5 | 0 | \( P^{(1)} \) |
| Dec 1999     | 1     | 2            | 0.3104        | 0 | 5/7 | 1/7 | 1/7 | 0 | 0 | \( P^{(2)} \) |
| Nov 1999     | 2     | 3            | 0.0967        | 0 | 4/13 | 0 | 2/13 | 5/13 | 2/13 | \( P^{(3)} \) |
| Oct 1999     | 2     | 4            | 0.0697        | 0 | 0 | 4/13 | 5/13 | 3/13 | 1/13 | \( P^{(4)} \) |
| Sept 1999    | 3     | 5            | 0.0810        | 2/29 | 5/29 | 4/29 | 10/29 | 8/29 | 0 | \( P^{(5)} \) |
| Aug 1999     | 4     | 6            | 0.0852        | 4/21 | 4/21 | 5/21 | 2/21 | 5/21 | 1/21 | \( P^{(6)} \) |
|              |       |              | \( P_i \) (weighted average) | 0.0337 | 0.4007 | 0.2162 | 0.1578 | 0.1673 | 0.0243 |

### Table 5. Stationary distribution and recurrence period of various states

| \( T_{ij} \) | \( \mathbf{z} \) | \( \mathbf{\mu} \) |
|------------|---------|---------|
| State(j)   | 1       | 2       | 3       | 4       | 5       | 6       |
| \( \pi_j \) | 0.0533  | 0.1333  | 0.2417  | 0.2000  | 0.2917  | 0.0750  |
| \( T_{ij}, \mu_j \) | 17.14 | 7.5 | 4.14 | 5 | 3.43 | 13.33 |

**CONCLUDING REMARKS**

The mathematical statistics tool is an important method for the prediction and forecast of infectious diseases. Historically, forecasting methods such as multivariate statistics analysis, Monte-Carlo simulations, spectrum analysis, that rely heavily on historical data have been used to infer future trends. But the accuracy of these non-subjective forecasting methods needs much improvement. In relation to these non-subjective forecasting methods, the weighted Markov chain theory introduced in this paper has the follow distinguishing characteristics:

① The key to the success of the forecast based on the weighted Markov chain theory in this article
is the scientific classification, determination of the initial state of the system, and the ensuring of the state transition forecasting methods have been heavily reliant on historical data, and largely affected by differences between historical and future environments.

② Since the weighted Markov chain is weighted with autocorrelation coefficient of various steps, the sum of the chain can be used to forecast the number of the infected. Therefore, it is more reasonable and sufficient in using data, and the Markov chain theory and the related analysis are well integrated. In the meantime, to calculate the limit distribution of the sequence applying the ergodic theorem reflects much more information of the sequence of the incidents of the disease in order to make a much more qualitative and quantitative description of the sequence calculated.

③ To determine the classifying criteria applying the ordered cluster, the data structure of the sequence of the patients can be taken full account of in the weighted Markov chain model, and the increase and decline in the historical data will be fully reflected. In this way, we are able to describe the status of the disease more accurately, so as to describe the internal distribution in a more effective way. Various methods in the multivariate statistics and the theory of fuzzy mathematics can be used to classify the state of the samples. The appliers should have a good understanding of the characteristics of the actual data, and accumulate experience in order to find more suitable classifying criteria.

④ With the continual increase of time sequence length, the representativeness of the historical data will be increased accordingly. The autocorrelation coefficient, transition probability matrix and the weight of various steps will change too, and this kind of change is also the process of improvement of the forecast and analysis theory. The forecasting model is not fixed, so the real number of the patients in every period of time should be added to the sequence of historical data. Therefore, the autocorrelation coefficient, transition probability matrix and the weight of the forecast can be adjusted online, and the accuracy of the forecast and analysis will be further improved. Moreover, the epidemic report of the disease forecast should have the same criteria in order to minimize the error and failure of reporting, and the disease information should be accumulated in the real practice.

⑤ With the development of the omy and culture, the improvement of hygiene conditions, and the strengthening of the prevention and control of epidemic diseases by the government, the epidemic diseases are controlled effectively, and the number of patients is declining year after year in China. In determining the structure of the model, all these changes should be paid attention to in order to make the statistical model more consistent with the life environment. Furthermore, as the number of the patients is able to reflect the change of the population and developing trend of the disease when the total population does not fluctuate too much, the paper applies the number of the patients to predict the future condition of the incidents of Hepatitis B in the coming year.

⑥ This forecasting method is effective when the spread and the prevention and control measures have not changed fundamentally. However, if preconditions are not met, the forecast will lose its value. Meanwhile, it is still challenging to calculate the actual number of the incidents of patients based on the state percentage calculated. It is very practical to see the occurrence and development of an epidemic disease as a stochastic process. The forecast and analysis method put forward in this article organically combines stochastic process theory, correlative analysis, ordered cluster analysis and epidemiology. Using an easy calculation and clear concepts, it provides a very good way to explore and discuss the forecast and prediction of epidemic diseases.

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