

Research Article

Spatial Cluster Analysis by the Adleman-Lipton DNA Computing Model and Flexible Grids

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Spatial cluster analysis is an important data-mining task. Typical techniques include CLARANS, density- and gravity-based clustering, and other algorithms based on traditional von Neumann's computing architecture. The purpose of this paper is to propose a technique for spatial cluster analysis based on DNA computing and a grid technique. We will adopt the Adleman-Lipton model and then design a flexible grid algorithm. Examples are given to show the effect of the algorithm. The new clustering technique provides an alternative for traditional cluster analysis.

1. Introduction

Deoxyribonucleic acid computing, or DNA computing in short, has attracted strong interests and wide focus recently. It is inspired by the similarity between the way DNA stores and manipulates information with traditional Turing machine. Although DNA computing is in a sense similar to evolutionary computing, but the significant difference between them lies in the computing medium, biomolecules rather than transistor chips. It is this difference that makes DNA computing a promising field with ultimate goal of making DNA computers [1].

The essential work to reveal the ability of DNA in computing is by Adleman's experiment (Adleman [2]), which demonstrated that the tools of laboratory molecular biology could be used to solve computation problems. Adleman also proves the huge information storage capacity of DNA which is contained in the sequence of nucleotide bases that hydrogen bonds in a complementary fashion to form double-stranded molecules from single-stranded oligonucleotides. Adleman's work was later generalized by Lipton [3] to the satisfiability problem. Based on Adleman and Lipton's research, a number of applications of DNA computing in solving combinatorially complex problems such as factorization, graph theory, control, and nanostructures have emerged [1]. There appeared also theoretical studies

including DNA computers which are programmable, autonomous computing machines with hardware in biological molecules mode, see [4–7] for references.

Adleman and Lipton's original works include a basic computing model, often referred to as the Adleman-Lipton model. Later generalizations include the sticker model, the splicing model, and the insertion deletion model [1]. However, most applications in this area are restricted to problems of combinatory types due to searching nature of DNA computing. It is a challenge how to design applications of optimization types.

Spatial cluster analysis is a traditional problem in knowledge discovery from databases [8]. It has wide applications since increasingly large amounts of data obtained from satellite images, X-ray crystallography, or other automatic equipment are stored in spatial databases. The most classical spatial clustering technique is due to Ng and Han [9] who developed a variant PAM algorithm called CLARANS, while new techniques are proposed continuously in the literature aiming to reduce the time complexity, or to fit for more complicated cluster shapes.

For example, Bouguila [10] proposed some model-based methods for unsupervised discrete feature selection. Wang et al. [11] developed techniques to detect clusters with irregular boundaries by a minimum spanning tree-based clustering algorithms. By using an efficient implementation of the cut and the cycle property of the minimum spanning trees, they obtain a performance better than $O(N^2)$. In another paper, Wang and Huang [12] developed a new density-based clustering framework by a level set approach. By a valley seeking method data points are grouped into corresponding clusters.

Although DNA computing and cluster analysis receive much attention and rapid development, there have appeared rare combination of these two important research areas. Up to the authors knowledge, the combination of DNA computing and cluster analysis is found in a few researches such as Bakar et al. [7].

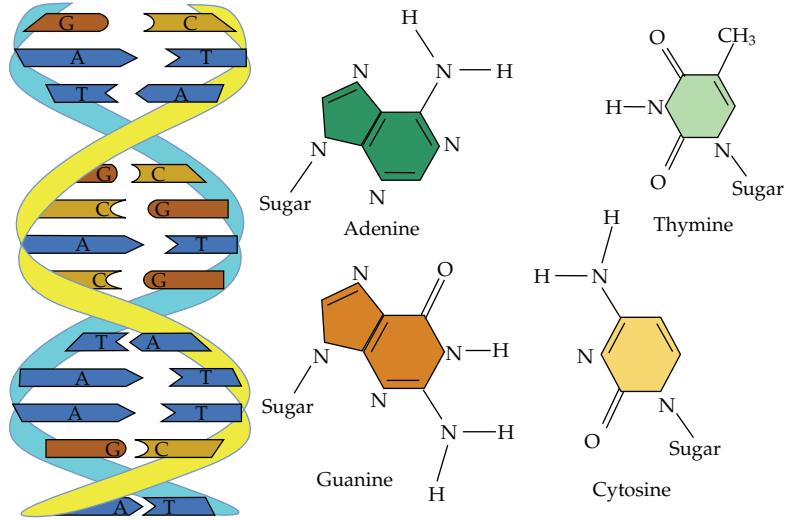
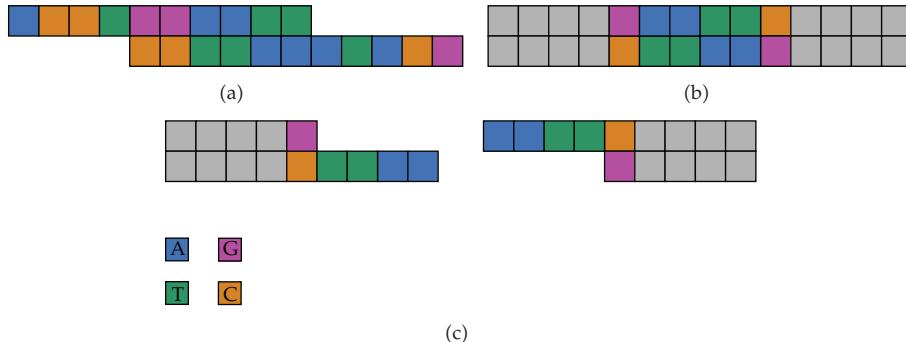
Inspired by the research of Bakar et al. [7], this paper focuses on the joint study of DNA computing with cluster analysis. We propose a new grid-based clustering technique which can be solved by DNA computing. Different with other researches, this can reduce the searching space significantly. Finally we present two examples to show the details of our technique.

2. DNA Computing and Operations

2.1. DNA Structures

Macromolecules of nucleic acids are composed of nucleotide building blocks. In DNA, the nucleotides are the purines adenine (A), guanine (G), the pyrimidines thymine (T), and cytosine (C). Single-stranded DNA molecules, or oligonucleotides, are formed by connecting the nucleotides together with phosphodiester bonds. The single strands of DNA can form a double-stranded molecule when the nucleotides hydrogen bonds to their Watson-Crick complements, $\bar{A} = T$ and $\bar{G} = C$ (Figure 1).

DNA stores information in nucleic acid and manipulates information via enzymes and interactions. A strand of DNA is encoded with four bases represented by the letters A, T, C, and G. Each strand has a 3'- and a 5'-end, and hence any single strand has a natural orientation. The cutting of certain strands of a DNA molecule is performed by the restriction enzymes. These enzymes catalyse the cutting operations at very specific DNA base sequences which are called recognition sites. Figure 2(a) shows a DNA molecule in which its four

**Figure 1:** A double-stranded DNA structure diagram.**Figure 2:** DNA molecules with sticky ends.

nucleotides in the left end and five in the right end are not paired with nucleotides from the opposite strand caused by cutting, or some other operations. In this case, the molecule is called to have sticky ends.

Here is an example which illustrates the process by the enzyme EcoRI as shown in Figure 2(b) where N represents some other arbitrary deoxyribonucleotide. EcoRI acts only at the six-term sequences which are exactly like the form

$$\begin{array}{ccccccc} G & A & A & T & T & C \\ C & T & T & A & A & G. \end{array} \quad (2.1)$$

The effect is to cut the molecule into two pieces as shown in Figure 2(c).

There are over 100 different restriction enzymes, each of which cuts at its specific recognition site. A restriction enzyme cuts DNA into pieces with sticky ends. On the other hand, sticky ends will match and attach to other sticky ends of any other DNA that has been

cut with the same enzyme. DNA ligase joins the matching sticky ends of the DNA pieces from different sources that have been cut by the same restriction enzyme.

2.2. DNA-Computing Models

There are several types of DNA-computing models among which the Adleman-Lipton model is the most traditional one. This model focuses on the hybridization between different DNA molecules as a basic step of computations. According to Adleman and Lipton's original works [2, 3], this traditional DNA-computing strategy is based on enumerating all candidate solutions, and then using some selection process to choose the correct DNA. This technique requires that the size of the initial data pool increases exponentially with the number of variables in the calculation.

Apart from the Adleman-Lipton model, other DNA computing models appeared such as the sticker model, the splicing model. The Sticker model is based on a coding scheme called DNA complex. A DNA complex is a partially double DNA strand. Usually a double piece represents a bit with value one while a single-strand represents zero. Hence each complex is constructed by two single stranded DNA molecules referred to as memory strands and sticker strands. A memory strand contains n nonoverlapping substrands each of which is m bases long. Each sticker strand is m bases long and is complementary to exactly one of the n substrands in a memory strand.

The second model is the splicing model proposed by Tom [6] based on formal language theory. A splicing system $S = (A, L, B, C)$ consists of a finite alphabet A , a finite set I of initial strings in A^* (language over A), and finite sets B and C of triples (c, x, d) with $c, d, x \in A^*$. Each such triple in B or C is called a pattern. For each such triple the string $cx d$ is called a site, and the string x is called a crossing. Patterns in B are called left patterns, and patterns in C are called right patterns. The language $L = L(S)$ generated by S consists of the strings in I and all strings that can be obtained by adjoining to $Lucxfq$ and $pexdv$ whenever $ucsdv$ and $pexfq$ are in L and (c, x, d) and (e, x, f) are patterns of the same hand. A language L is a splicing language if there exists a splicing system S for which $L = L(S)$.

The next model is the k -armed model which is based on some more complicated molecule structures which have three-dimensional DNA architecture. In [13] the authors pointed out that it is natural to use the armed model to represent SAT problem in terms of contact network framework, and they gave theoretical solutions to this NP-complete problems. Like the splicing model, biological operations in the k -armed model include cleaving and connecting.

2.3. Operations of the Adleman-Lipton Model

The basic principle of DNA computing is to use the encoded information in the sequence of nucleotides and evolve them by breaking and making new bonds between them to reach the answer. The basic operations performed by enzymes are denaturing, replicating, merging, detecting, and so forth.

According to the DNA computing models proposed by Adleman [2] and Lipton [3], there are several basic DNA operations. One important operation is hybridization which is a main process in DNA computing to form all possibilities of solution strands in which the right answer lies. Hybridization is done by mixing strands in tubes with the help of some enzymes.

Apart from hybridization, the basic DNA operations available on DNA are mainly the following.

- (i) Merge. $m(N_1, N_2) \triangleq N_1 \cup N_2 = N$.
- (ii) Amplify. $Duplicate(N_1) = N$.
- (iii) Detect (N).
- (iv) Separate or extract. $N \leftarrow +(N, w)$, $N \leftarrow -(N, w)$. Given a word w consisting of strings from $\Sigma = \{A, G, C, T\}$ and a tube N , generate two tubes $+(N, w)$ and $-(N, w)$ which contain and does not contain the string w .
- (v) Length separate. $N \leftarrow (N, \leq n)$. Given a tube N and an integer n , generate a tube containing stands with length less or equal to n .
- (vi) Position separate. $N \leftarrow B(N_1, w)$, $N \leftarrow E(N_1, w)$. Given a tube and word generate a tube with stands beginning (ending) with the word.

3. Grid-Based Clustering

The grid-based clustering uses a multiresolution grid structure, called cells, which contains the data objects and acts as operands of clustering performance. Traditional approaches include STRING WaveCluster, and CLIQUE [8]. The most common grids are regular hypercubic grid. This requires that the grid construction covers all the data space with the same precision. The second method uses flexible grids, that is, multiresolution grids with hypercubic or hyperrectangular cells having randomly oriented borders [14].

3.1. A Flexible Grid Definition

Suppose that the data set is $\Omega = \{x_1, \dots, x_N\} \subset R^n$. It is bounded by a rectangle D_0 in R^n . A grid is an undirected graph $G = (V, E)$ where each node of V is called a cell and is represented by a quad $v = (D, c, p, \sigma)$, where D is a polyhedra, c is a center point of D , $p = |\omega \cap D|$ is the number of points of Ω covered by the cell, and σ is the diameter of D . We will always assume that a cell is nontrivial; that is, D has interior points in R^n . For a cell $v = (D, c, p, \sigma)$, its boundary is denoted by ∂D which is the set of hyperplane pieces bounding the polyhedra. If $S \in \partial D$ and S is part of a hyperplane H , then H is called a *tangent plane* of the cell. Two nodes $v_i = (D_i, c_i, p_i, \sigma_i)$, $i = 1, 2$, are called *adjacent* if the two cells share a common tangent plane and $D_1 \cap D_2 \neq \emptyset$. For two adjacent cells, define an edge between them. Hence, $E = \{uv : u$ and v are adjacent $\}$. Figure 3 presents an illustration of tangent plane and adjacent cells.

To construct the graph G , we need two parameters p_0 and σ_0 indicating the minimum number of points to be considered, and the minimum diameter. Then the graph is constructed iteratively. We start with the first node $v_1 = (D_0, c_1, N, \sigma_1)$, where D_0 is the original rectangle, c_0 is the center of D_0 . Then at each step, the cell containing dense points (controlled by a threshold value p_0), or with larger diameter (controlled by threshold value σ_0), is split into two subcells by a hyperplane. A cell is sparse if it contains less points than p_0 . It is called a small cell if its diameter is less than σ_0 . If we reach a sparse or small cell, then add this cell to the node set of the graph. This step continues until no more cell left to be split. The resulting graph is called a flexible grid. Algorithm 1 gives the algorithm for the graph construction process.

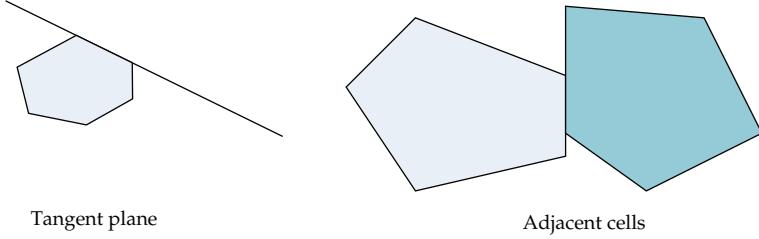


Figure 3: Illustration of tangent plane and adjacent cells.

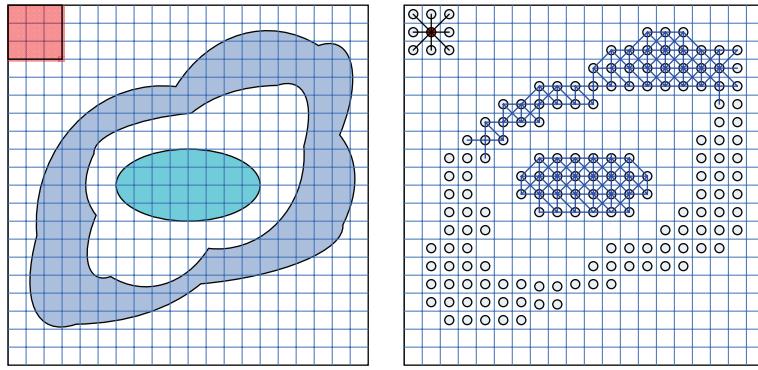


Figure 4: An example of flexible grid with induced graph.

We present an example to show the data set and the flexible grid generated by the above algorithm (Figure 4).

Next we define the weights on edges. A weight on an edge is the dissimilarity of the adjacent nodes. Suppose that the Euclidean distance in R^n is denoted by $d(\cdot, \cdot)$. Here and after, we will always assume that a data point $x \in D$ means $x \in D \cap \Omega$. Then for two nodes $v_i = (D_i, c_i, p_i, \sigma_i)$, $i = 1, 2$, the weight is defined by

$$\omega(v_1, v_2) = \begin{cases} 0, & \text{if } \min\{p_1, p_2\} = 0 \\ \frac{1}{p_1 p_2} \min_{x \in D_1, y \in D_2} d(x, y) & \text{if } \min\{p_1, p_2\} > 0. \end{cases} \quad (3.1)$$

3.2. Clustering Problems

When the graph is constructed, the clustering problem is converted into grouping nodes of the graph into clusters. Traditional techniques include the hierarchical clustering [8]. For the purpose of this paper, we will give a different approach for this problem. First it should be noted that nodes corresponding to sparse areas are outliers. Therefore, in order to reduce computing complexity, we first remove all sparse graph nodes with corresponding edges. We still use $G = (V, E)$ to denote the resulting graph, where V is the set of vertices, and E the set of weighted edges. An example is shown in Figure 4 with part of its edges.

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Inputs:  $\Omega = \{x_1, x_2, \dots, x_N\}$  dataset of  $N$  points in  $R^n$ ,  $D$ : hyper-rectangle containing  $\Omega$ ,  $p_0$ :  

    population threshold value,  $\sigma_0$ : cell diameter threshold value
Outputs:  $V = \{v_1, v_2, \dots\}$ : set of vertices
Begin
 $V(1) = \{v_1 = (D_0, c_1, N, \sigma_1)\}$ . Let  $t = 1$  and  $D_1 = D_0$ .
while  $V(t) \neq \emptyset$  do
    for each  $v = (D, c, p, \sigma) \in V(t)$  do
        Choose a cutting hyperplane  $L$  passing  $c$ . Cut the current cell  $v$  into two subcells  $v', v''$ .
        For each  $v \in \{v', v''\}$  of the new cells, if  $p_v < p_0$  or  $\text{diam}(D_v) < \sigma_0$ , add this new cell  $v$  to the node set  $V$ . Else add  $v$  to  $V(t + 1)$ .
    endfor
     $t = t + 1$ ;
end
End

```

Algorithm 1: A flexible grid construction algorithm.

Now we consider the problem of weight computation. By the graph construction procedure, we know that any node will correspond to a cell with diameter no larger than σ_0 . Therefore, the distance between cells can be approximated by $d(c_1, c_2)$, and (3.1) will be

$$\omega(v_1, v_2) = \frac{1}{p_1 p_2} d(c_1, c_2). \quad (3.2)$$

In this way we can significantly reduce the computing time without loss of much precision. Again we define a parameter $0 < \omega_0 \leq \infty$. We will eliminate those edges with weight $\omega > \omega_0$. If this parameter $\omega_0 = \infty$, this means that no edges are eliminated. Now we use $\mathcal{C} = \{V_q : q = 1, 2, \dots, k\}$ to denote a clustering of the vertices set V of graph G for threshold values p_0 and σ_0 . Next we will use $|V_q|$ to denote the number of its vertices. Define the energy of clustering as follows:

$$\begin{aligned} \min \mathcal{E}(\mathcal{C}) : \mathcal{E}(\mathcal{C}) &= \sum_{q=1}^k \sum_{\substack{v_i \neq v_j \\ v_i, v_j \in V_q}} \omega(v_i, v_j), \\ \min \mathcal{E}_{\text{int}}(\mathcal{C}) : \mathcal{E}_{\text{int}}(\mathcal{C}) &= \left(\min_{\substack{1 \leq i, j \leq k \\ i \neq j}} \min_{\substack{u \in V_i \\ v \in V_j}} \omega(u, v) \right)^{-1}. \end{aligned} \quad (3.3)$$

Then the clustering problem is a minimization of the energy functions. However, the optimization problem is hard to solve. We will present a variation in the following.

3.3. Path Clustering

Now we consider the graph $G = (V, E)$ with weight matrix W . Assume that the number k of clusters is a positive integer. A Hamiltonian path \mathcal{L} of G is a path that visited each vertex exactly once. Now we remove $k - 1$ nonadjacent edges from \mathcal{L} and denote the result by \mathcal{L}_k .

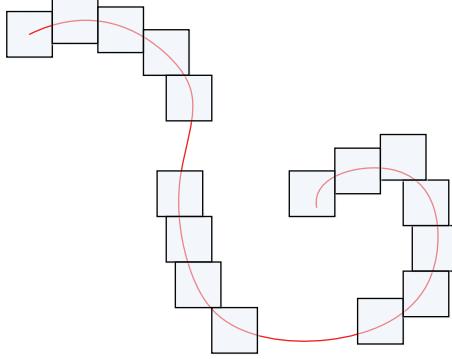


Figure 5: An example of path clustering with three clusters.

Define the energy $E(\mathcal{L}_k)$ of L_k as the sum of its edge weights. The path with least energy $E(\mathcal{L}_k)$ is called a path clustering

$$\min \begin{cases} E(\mathcal{L}_k) : & \text{for any } \mathcal{L}. \\ E(\mathcal{L}_k) : & \text{for any } \mathcal{L}, 2 \leq k \leq N. \end{cases} \quad (3.4)$$

The first minimization of (3.4) is clustering for a fixed k , and the latter is clustering without fixing k . Path clustering is slightly different with distance-based clustering. The next example illustrates path clustering (Figure 5).

4. Clustering by DNA Computing

In this section we consider clustering of the graph $G = (V, E)$. We still use N to denote the number of nodes in V , and this will not cause any confusion. Suppose that the number of clusters is k which is *a priori* determined, or defined in the process of clustering. The problem is to partition the vertex set V into k clusters. Suppose that the original data set Ω is bounded by a constant $M/2 > 0$, that is, $\|x\| \leq M/2$ for $x \in \Omega$. Here we use the Euclidean distance for points in R^n and $\|x\| = \sqrt{a_1^2 + \dots + a_n^2}$, where $x = [a_1, \dots, a_n]$. Points in Ω are denoted by x_i and $\Omega = \{x_1, \dots, x_N\}$. A point in the data set Ω will be denoted by the lower case letter x .

For each point $x \in \Omega$ and a cluster C , define the distance between them as $d(x, C) = \min_{z \in C} d(c, z)$ where $d(\cdot, \cdot)$ is the Euclidean distance. If C_1, C_2 are two clusters, then define $d(C_1, C_2) = \min_{x \in C_1} \min_{y \in C_2} d(x, y)$. Clearly these distances are bounded by the constant M . For $u, v \in V$ define the dissimilarity measure as $\rho(u, v) = w(x, y)/M$. Clearly these dissimilarity measures are in the interval $[0, 1]$.

Now we convert the dissimilarity measures into integers. First we need to define an acceptable error rate $\varepsilon > 0$. This means that we do not distinguish those measures where their difference is less than ε . Now we divide the interval $[0, 1]$ into I subintervals with equal width $I^{-1} < \varepsilon$. For $z \in [0, 1]$ let its corresponding integer be $s(z) = [Iz]$ where operator $[\cdot]$ is the largest integer without exceeding it. Hence the dissimilarity measure lies in the set $\{1, \dots, I\}$.

Now we define the weight matrix on the graph \mathcal{G} by

$$W = [w_{ij}]_{N \times N}, \quad w_{ij} = s(\rho(x_i, x_j)), \quad i, j = 1, \dots, N. \quad (4.1)$$

A clustering of the graph \mathcal{G} , denoted by \mathcal{C} , is a partition $V = \cup_{i=1}^k C_i$. Thus any clustering can be taken as a rearrangement of the vertices v_1, \dots, v_N . For example, the vertex set $\{\{v_3, v_2, v_1\}, \{v_5, v_4\}, \{v_6, v_7, v_8, v_9\}\}$ with three clusters can be written as

$$\mathcal{C} = v_3 v_2 v_1 \alpha v_5 v_4 \alpha v_6 v_7 v_8 v_9. \quad (4.2)$$

Here we use the greed letter α as a separator between clusters. Therefore, we have $k - 1$ separators if we obtain k clusters. If we take dissimilarity measure into account, then a clustering will be as follows:

$$\mathcal{C} = v_3 w_{32} v_2 w_{21} v_1 \alpha v_5 w_{54} v_4 \alpha v_6 w_{67} v_7 w_{78} v_8 w_{89} v_9. \quad (4.3)$$

Now we have converted the clustering problem to a permutation problem. That is, any permutation of the set $\{1, 2, \dots, N, \alpha, \dots, \alpha\}$ (the number of α 's is $k - 1$) is a candidate solution. The string with minimum length is the optimal solution.

4.1. DNA Coding for Data

First we give the encoding of the dissimilarity measure, or the weights. In order to do this, we assume that 1 is coded by the string AG . And any integer $i \in \{1, \dots, I\}$ is coded by $\text{seq}(i)$:

$$\text{seq}(i) = \overbrace{AGAG \cdots \cdots AG}^{i \text{ number of } AGs}. \quad (4.4)$$

Next we present the encoding of a vertex $v \in V$. This is done by using a fixed mer sequence as the following example (20-mer):

$$\text{seq}(v) = 5' - TCTCT \ CTCTC \ TCTCT \ CTCTC \ TCTCT - 3'. \quad (4.5)$$

The separator α is also coded with a single-strand $\text{seq}(\alpha)$ of the same length with that of points. This is done exactly as the data point coding except that we need to distinguish separators with data points.

4.2. Encoding Scheme

Now we need to put everything together for a candidate solution (4.3). First we design a code for an edge $uv \in E$ with weight w . This is done by linking the last half part of $\text{seq}(u)$, $\text{seq}(w)$, and the first half of $\text{seq}(v)$ as shown in Figure 6(a). We need two special edges called the left half-edge and the right half-edge. The left half edge is a linking of $\text{seq}(u)$, $\text{seq}(w)$,

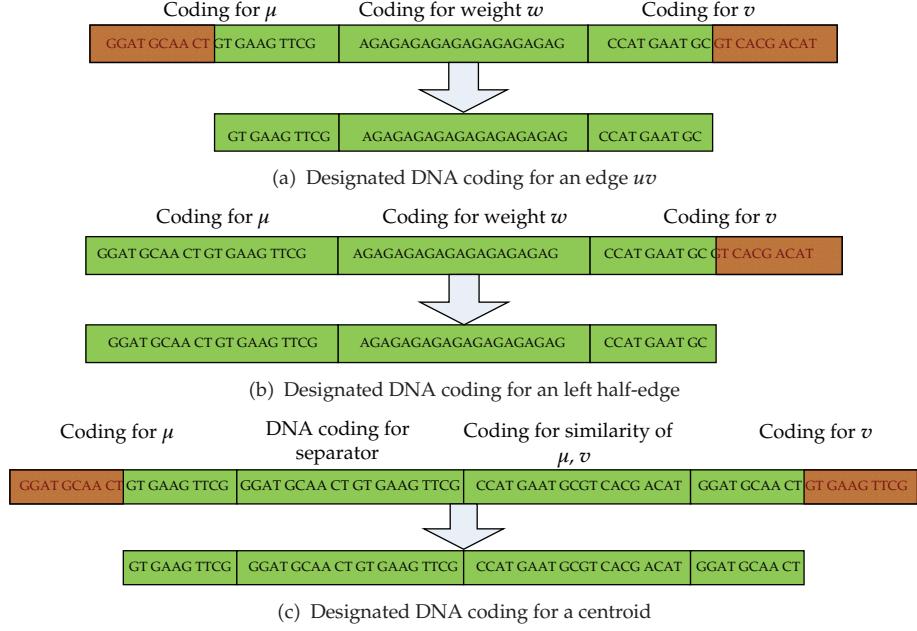


Figure 6: DNA-encoding architecture.

and the first half of $\text{seq}(v)$. The right half edge is a linking of the first half of $\text{seq}(u)$, $\text{seq}(w)$, and $\text{seq}(v)$.

Next we define coding for a centroid which is a complementary form of a separator α . First for two vertices $u, v \in V$, we define its similarity as $\hat{\rho}(u, v) = I - s(\rho(u, v))$. One should notice the difference with dissimilarity measure defined in the matrix W . Now we define the code of a centroid as a linking of the last half part of $\text{seq}(u)$, $\text{seq}(\alpha)$, $\text{seq}(\hat{\rho}(u, v))$, and the first half of $\text{seq}(v)$ as shown in Figure 6(c). Here u, v are two vertices in V .

Finally, the code for a cluster C is a permutation of the following string without changing the position of the left half-edge and the right half-edge:

$$\text{seq}(C) = \underbrace{A \cdots C}_{\text{left half-edge}} \underbrace{\overbrace{A \cdots C}^{\text{edges}} \overbrace{G \cdots T}^{\text{centroid}}}_{N-k-1 \text{ edges}, k-1 \text{ centroids}} \underbrace{\overbrace{G \cdots T}^{\text{centroid}} \overbrace{A \cdots C}^{\text{edges}}}_{\text{centroids, edges}} \underbrace{T \cdots G}_{\text{right half-edge}}, \quad (4.6)$$

N points and $k-1$ separators altogether

4.3. DNA Program

Now we describe the biological operations for the clustering. First we put single strands in a tube T_0 including complementary strands of all vertices, complementary strands of the separator α , and complementary strands of integers $1, 2, \dots, I$. These strands serve as splints. We also pour strands into T_0 of all the left half-edges, right half-edges, strands of integers $1, 2, \dots, I$, all edges, and all centroids. Then hybridization and ligation can be executed. As a result, all combinations representing clustering schemes are obtained in the tube T_0 .

Step and Procedure

- (1) **Input** (T_0).
All DNA sequences and complementary DNA sequences are placed in empty test tube T_0 .
 - (2) **Amplify** (T_0).
Make all sequences in T_0 mixed together and execute ligation process. After hybridization process, all possible combinations of DNA sequences happen in T_0 .
 - (3) $T_1 \leftarrow +(T_0, \text{seq}(\alpha))$.
Select only those DNA strands which include at least one separator α from T_0 and keep them into empty test tube T_1 .
 - (4) **for** $i = 1$ to N **do** {**begin** $T_1 \leftarrow +(T_1, \text{seq}(v_i))$ **end**; }**endfor**; $T_2 \leftarrow T_1$.
Select all DNA strands that contain all the N vertices v_1, \dots, v_N in test tube T_1 . Put them in empty test tube T_2 .
 - (5) **Gel Electrophoresis**.
Find the shortest DNA sequence in test tube T_2 . Put them in an empty tube T_3 . This is the solution of clustering the problem.
 - (6) **Count the number of separators** α .
Amplify and count the number of clusters in tube T_3 .
- END

Algorithm 2: The DNA program.

In order to select acceptable solutions among these combinations, we need to eliminate those strands which do not contain a separator α and those which do not contain all the vertices v_1, \dots, v_N . Finally, by counting the number of separators α as k of the strand with shortest length, we get the solution of the clustering problem. This final procedure can be implemented by direct observation to calculate the separator sequences by using special microscope such as atomic force microscope (AFM) to identify and calculate marking sequences [7].

The DNA program of biological operations is shown in Algorithm 2.

5. Examples and Discussions

In this section, we present some examples to illustrate the performance of our algorithm. Then we will show that our technique will give clusters more naturally.

5.1. Example One

First we present an example with 20 points to be clustered that is discussed in [7]. First we construct a grid with interval 1 as shown in Figure 7. Then we take $p_0 = 1$ as minimum points located in each grid. As a result, the induced graph is disconnected with 10 subgraphs. The final result shows four clusters with additional 6 outliers. This is different with result of [7] where they obtain four clusters with no outliers.

Now we change the grid into interval 2. Then we have only half the grids compared to grid with interval 1. We take $p_0 = 1$ but we use another parameter $\omega_0 = 1$. This time we surprisingly obtain three clusters (Figure 8) with two outliers. This fact shows that the clustering method proposed here is sensitive to the construction of grids and parameters. Considering various definitions and measurements of clustering, this is not too surprising.

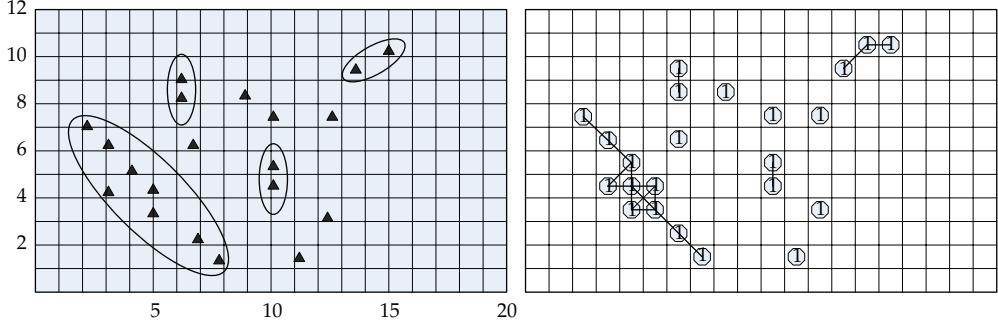


Figure 7: Data example one with graph generated with grid interval 1. Parameters $p_0 = 1$.

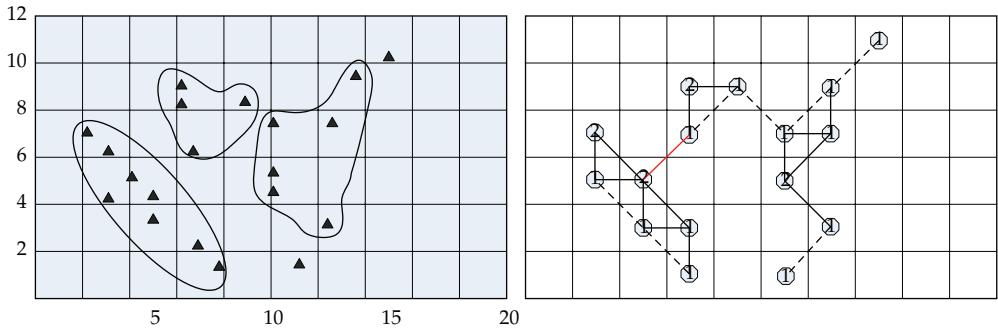


Figure 8: Data example one with graph generated with grid interval 2. Parameters $p_0 = 1$ and $\omega_0 = 1$.

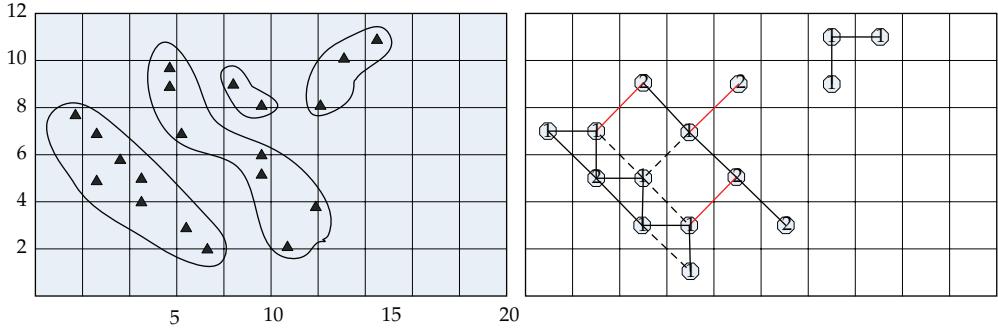


Figure 9: Data with different grid (interval 2). Parameters $p_0 = 1$ and $\omega_0 = 1$.

Now we construct a grid similar with Figure 8 but starting the first horizontal vertical lines not from coordinates $(0, 2)$ and $(2, 0)$. Identically, we can implement this case by moving the whole data set in the grid of Figure 8. With the same parameters as above, we obtain the third clustering result as shown in Figure 9. Now four clusters are generated without outliers. It is interesting to note that the three different grids induce one common cluster (left-bottom cluster). In fact, this common cluster is better organized than the other clusters. Hence we know that clustering is sensitive to the construction of grids especially for those *bad* clusters.

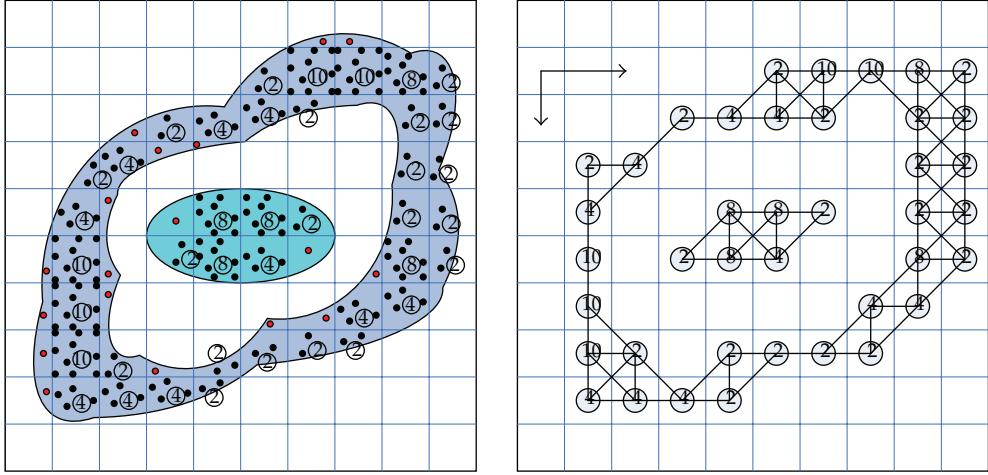


Figure 10: A data example with graph generated. The numbers in circles are the values of p in cells.

5.2. Example Two

Now we consider another example as shown in Figure 10 with the data to be clustered and the graph constructed. For this example, we take $p_0 = 2$. For adjacent cells, if they share a common edge, then we define their dissimilarity measure as 1. Otherwise if they share a common vertex, then define dissimilarity measure as 1.4. There are 40 nontrivial cells.

Since for any two cells $p_1 p_2 \leq 100$, we take $I = 100$ and by (4.1) the weight matrix is

$$W = \begin{bmatrix} W_1 & W_2 \\ W_3 & W_4 \end{bmatrix}. \quad (5.1)$$

Here W_1 , W_2 , W_3 , and W_4 are 20×20 matrices, and W is symmetric. The matrix W_1 , W_3 , W_4 is shown in Table 1. The weighted graph is shown in Figure 11.

By the technique of this paper, the solution of clustering is a string. One of the clustering string as a clustering is as follows:

$$\begin{aligned} & v_{17}v_{18}v_{19}v_{25}v_{24}v_{23}\alpha v_6v_7v_8v_2v_1v_9v_3v_4v_5v_{11}v_{10}v_{14}v_{15}v_{21}v_{20}, \\ & v_{26}v_{27}v_{30}v_{29}v_{36}v_{35}v_{34}v_{33}v_{40}v_{39}v_{38}v_{37}v_{31}v_{32}v_{28}v_{22}v_{16}v_{13}v_{12}. \end{aligned} \quad (5.2)$$

6. Clustering of the Iris Data

In this section, we present another detailed examples to illustrate the encoding of the DNA-based clustering technique proposed in previous sections. The new example is the well-known Iris flower data set problem. The Iris flower data set is introduced by Sir Ronald Aylmer Fisher as an example of discriminant analysis [15]. The data set consists of 50 samples from each of three species of Iris flowers, that is, Iris setosa, Iris virginica, and Iris versicolor. Four features were measured from each sample, and they are the length and the width of sepal and petal [15, 16].

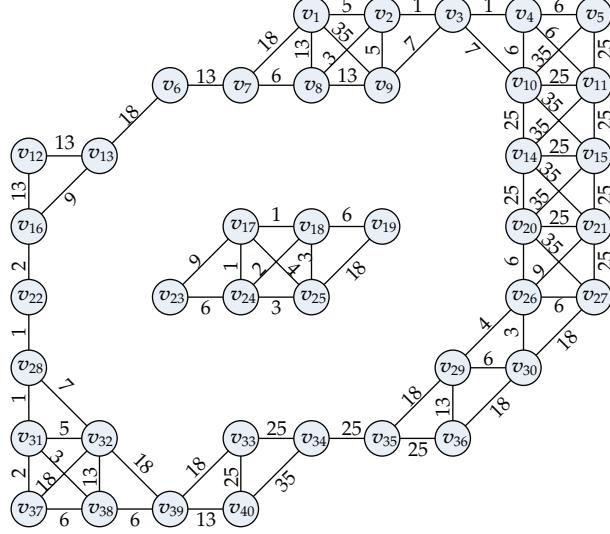


Figure 11: The constructed graph with weights on edges.

6.1. Grids and Graph of the Problem

Now we use a matrix $X|_{150 \times 4}$ to denote the data set. Then X is located in a rectangle $[4.3, 7.7] \times [2.0, 4.4] \times [1.1, 6.9] \times [0.1, 2.5]$ which is in the four-dimensional space R^4 . A grid cell (rectangle) D in R^4 has $2^4 + 4 \times 2^3 + 4 \times 2^2 + 4 \times 2$ adjacent cells. If we choose $D_0 = [4.01, 8.01] \times [1.81, 5.01] \times [1.01, 7.01] \times [0.01, 3.01]$ for instance, then the grids are defined as follows:

$$\begin{aligned} g_1 &= [4.01, 4.21, 4, 41, 4, 61, 4, 81, 5, 01, \dots, 7.81, 8.01], \\ g_2 &= [1.81, 2.01, 2.21, 2.41, 2.61, \dots, 4.81, 5.01], \\ g_3 &= [1.01, 1.21, 1.41, 1.61, 1.81, 2.01, \dots, 6.81, 7.01], \\ g_4 &= [0.01, 0.21, 0.41, 0.61, 0.81, \dots, 2.61, 2.81, 3.01]. \end{aligned} \tag{6.1}$$

Then the cells can be denoted by $\mathfrak{D} = \{D_{pqrs}|_{20 \times 16 \times 30 \times 15}\}$ with 144000 cells which is a huge number. Due to this reason, we choose $x_2 x_4$ among the four dimensions as shown in the first image in Figure 12 as cluster feature variables. The second figure in the same image shows the other two dimensions $x_1 x_2$ which is studied in Qu et al. [16].

Next we design a flexible grid structure as shown in Figure 13. Choose parameter $p_0 = 0$. Then the induced graph is shown in Figure 14. By (3.2) and direct computation, we obtain dissimilarities as matrices of the graph. Now we define the error rate $\varepsilon = 0.001$ and $I = 1000$. We cut the weight value by a maximum value 999. Then the weight matrices of G are shown as in Tables 2 and 3. Here we write the matrix as

$$W = \begin{bmatrix} W_{11} & W_{12} \\ W_{21} & W_{22} \end{bmatrix}, \quad W_{11}, W_{12}, W_{21}, W_{22} \text{ are } 13 \times 13 \text{ sub-matrices.} \tag{6.2}$$

Table 1: Sample data weight matrix [W_1, W_3, W_4].

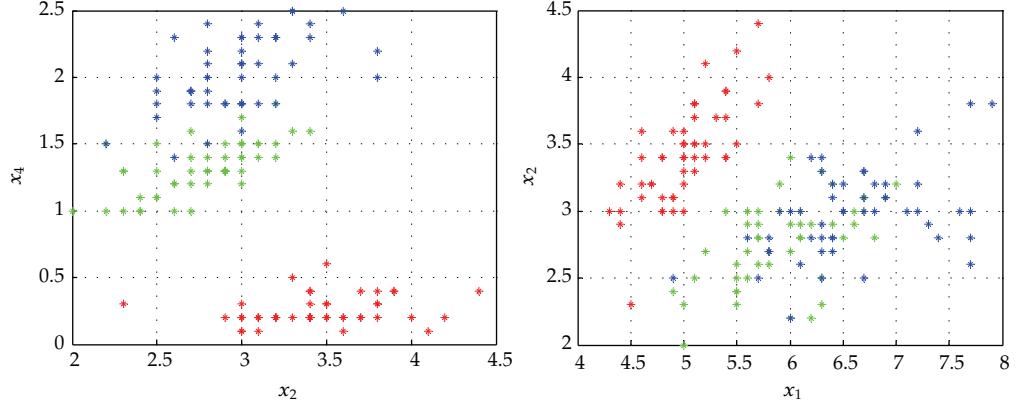


Figure 12: The Iris data figures for dimensions $x_2 - x_4$ and $x_1 - x_2$.

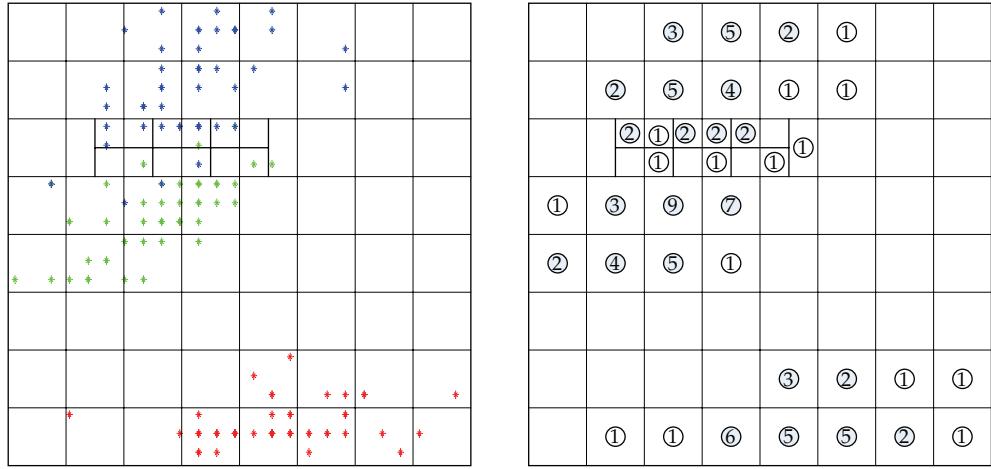


Figure 13: The Iris data figure for dimension $x_2 - x_4$ and with grids and candidate graph nodes.

For the two subgraphs as in Figure 14, we can find Hamiltonian paths easily. For subgraph two, the path is

$$v_1 v_2 v_7 v_8 v_{11} v_6 v_5 v_4 v_3 v_{10} v_9. \quad (6.3)$$

There exists many paths for subgraph one. One shortest path with indicator α along the edge $v_{16}v_{22}$ as shown in Figure 15(a) is

$$v_{11} v_3 v_2 v_1 v_4 v_5 v_6 v_{12} v_{13} v_{15} v_{18} v_{10} v_9 v_{17} v_8 v_{14} v_7 v_{16} v_{22} v_{23} v_{20} v_{26} v_{25} v_{21} v_{24} v_{19}. \quad (6.4)$$

The detailed encoding scheme is shown in the Appendix. The final clustering result is shown in Figure 15(b). The number of points which is not correctly clustered is 7. Clearly this is much better than other methods such as CEPSO of [16] where the error number is more than 20.

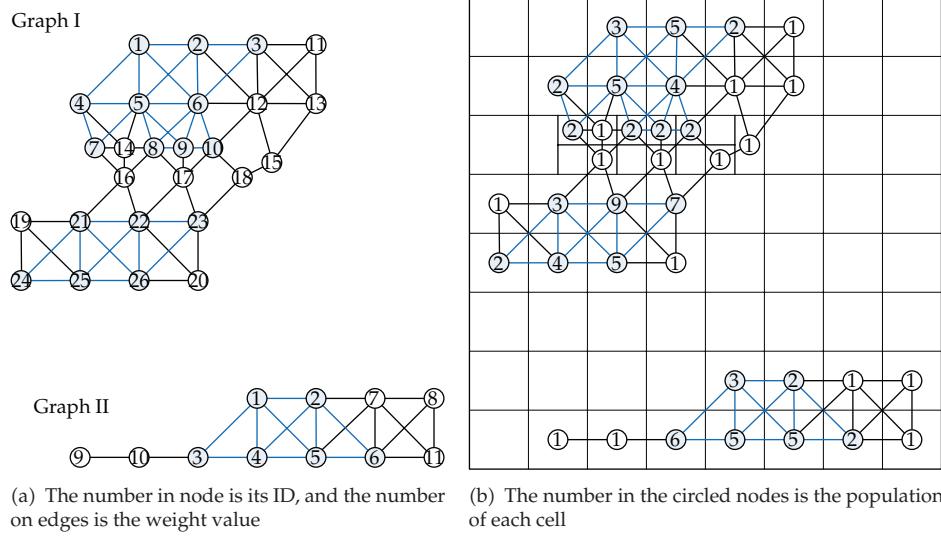


Figure 14: The induced graph with weight.

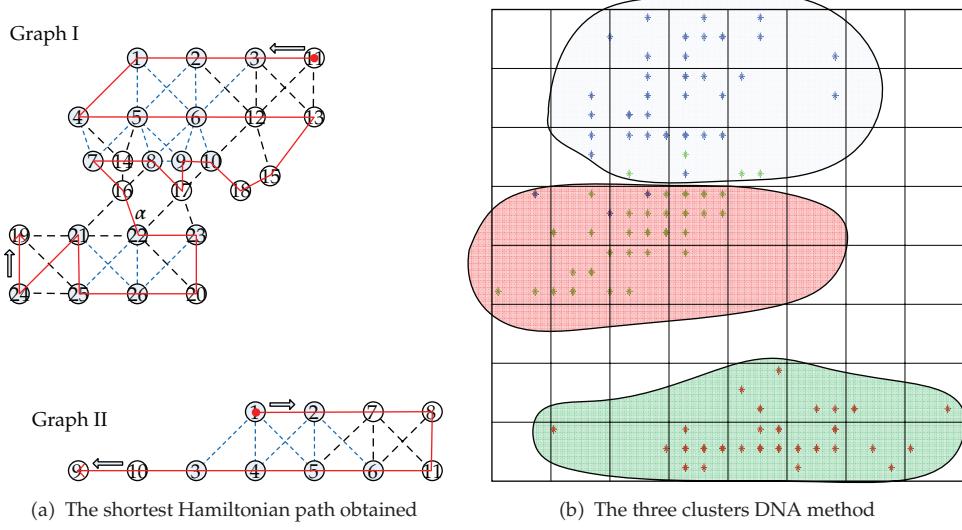


Figure 15: A short Hamiltonian path for the induced graph.

6.2. Discussions

Now we present some discussions about the time and computational costs of the proposed technique in this section. According to Algorithms 1 and 2, the computational costs consist of two main procedures, that is, the grid construction and biological operations, and we will show that the complexity is roughly linear.

First we check Algorithm 1 for the construction of flexible grids. The time complexity is the total searching counts $\sum_{t=1}^{t=T} |V(t)|$ where $|V(t)|$ is the cardinality of $V(t)$ and T is the final t that $V(t)$ reaches null. Clearly $|V(t)| \leq 2^t$. Now we estimate the upper bound of T .

Table 2: Dissimilarity weight matrices W_{11}, W_{12}, W_{22} of subgraphs I.

0	67	236	67	118					
	0	100		56	50				283
		0		177				500	500
			0	100		198			707
				0	50	106	79	106	
					0		133	99	99
						0			250
							0	125	
							0	125	
								0	395
								0	999
								0	999
									0
0									
	0								
		0							
			0						
530			0						
158				0					
					0				
250		354				0			
250		354	354				0		
			250				0		
			354	354				0	
								0	
								0	0
999									0
999									0
0	500								
	0		354						
		0			354				
			0			88			
				0		118			
					0		113		
						152			
							500		
								354	
									200
								0	
								37	
									143
									236
									83
									94
									0
									39
									22
									40
								0	
								125	
									0
									50
									0

In the worst case when $V(t)$ consists of 2^t cells, and each cell has a data population larger than p_0 , we have

$$p_0 2^t \leq N. \quad (6.5)$$

Table 3: Dissimilarity weight matrix of subgraphs II.

0	167	79	67	94				
	0		141	100	354	500		
		0	33				167	
			0	40				
				0	100	183		
					0	500	707	500
						0	999	999
							0	999
								0
								0

There we obtain the estimate $T \leq \log(N/p_0)$. And the total time upper bound is a linear time as follows:

$$\sum_{t=1}^T |V(t)| \leq \sum_{t=0}^T 2^T \leq 2^{T+1} \leq \frac{2N}{p_0}. \quad (6.6)$$

Next we analyze the complexity of DNA program as proposed in Algorithm 2. It is clear that the DNA program consists of three searching procedures. Step 3 selects those strands which contain at least one marker α which is a $O(1)$ operation. Step 4 checks if we get a sequence containing all the vertices with a complexity of $O(N_v)$, where N_v is the number of vertices and $N_v \leq N$. The final Step 6 counts the number of α 's and splits the sequence into clusters. For the shortest sequence which contains all the vertices, the maximum number of markers α is N_v . Hence the total DNA complexity of the program is $O(N)$ which is linear.

Finally we make some comments about experiments. By Adleman [2] and Păun et al. [1], when we design a DNA program for solving the problem, a test tube (as taken in the laboratory) is considered a multiset of words (finite strings) over the alphabet $\{A, C, G, T\}$ with basic operations proposed as in [1, 2]. These basic operations are standard and biologically implementable.

However, when we want to simulate the algorithm with personal computers, there will appear some tricky complexity that lay behind the biological operations. Again we analyze the steps in Algorithm 2. First we consider Step 2, and we only consider generating sequences containing each vertex exactly once. The time complexity here will be $O(2^{N_v}) = O(2^N)$. By examining other steps, we find the total time is $O(2^N)$.

7. Conclusion

In this paper we presented a new DNA-based technique for spatial cluster analysis. Two examples are given to show the effect of our algorithm. We do not need the number of clusters in advance. By a flexible grid method we can reduce the size of searching space significantly. This is different with other DNA-based applications in that they enumerate all solutions which will produce a large search space. This is especially useful for large database even through DNA computing that has large parallel ability. Also by changing the grid and corresponding parameters, we can get various clusters.

Finally we will point out that nonconvex clusters can be easily obtained by the technique of this paper. Comparing to research in [7] where only convex clusters can be generated, this is useful for complex database to be clustered. Up to the authors knowledge, this is, the first research in cluster analysis by DNA computing for nonconvex clusters. It provides an alternative solution for this traditional knowledge-engineering problem, which is *not* a combinatorial in nature. Comparing the many applications of DNA computing mainly in combinatorial problems, this is still interesting.

Appendix

DNA Encoding of the Iris Data

In this appendix, we present a DNA-coding scheme for the Iris data problem. By Tables 2 and 3 the weight values range from 16 to 999. Therefore, we need a maximum length of 999 mer to code the weights.

First we present one coding scheme for vertices of Subgraph I.

VERTEX CODING:

1. GGCCCATT GTCGACGA
2. AAGCATCC GAACGTCT
3. ATTTAGTA ACTGTCAT
4. CCACTAGG CATTACT
5. TAACTGAA TAGCCTAA
6. CCCATCCT GACTGCAT
7. GATGTGCT CGTAGCCT
8. AACCCCTA CGTGCATC
9. ATGCCCAA CATATAGG
10. TATTTAGT TGAAGCAT
11. TTCAGGAG GTGCACTA
12. CGTTTAGC CACCTTAG
13. TGTATACA CTTTATGT
14. CAACCATA GCCGCCGT
15. TCCCCGTC ATACCTTG
16. TAGGTATG ACTCGGCG
17. AGAAAATCG GCCGCAAT
18. GAGCGACT TCCTGGGT
19. CAATCATT TGCGGAGT
20. GCCGTGAT ACCCTCAG
21. AATGCACA GAGTTAA
22. TAGACTGT GTAAATCG
23. TCTACCGC CGTCCTTG
24. AATGAACA TTAGTGGC
25. TCTGCATA CGCCCCGT
26. AGCGTTGT GTCCTTCT

Since the edge coding of $v_i v_i$ is similar to those of $v_i v_j$, we next only present coding for one edge between each two vertices as follows.

EDGE CODING:

1-2: CAGCTCGT GGCCCAACGCCATTCTGTACCAACGAGAACCAAATAACGGAGC-
 ATAGAAGGGCATCTTACTTGGAG TTGCTAGC 1-4: CAGCTCGT AGCTACACGAGC-
 TTCCGTATCCCTGTGCGAATGCTGGCCTCTGCGGAGTTACGGGG CCCCTCAG-
 ACCCAGAACGACTGTTATGCTGCCAGCTGTCAATGAATCTTAGAGTGTG AACG-
 ACATATCCTGCGATTGTAG ATTGTTATCCTGAAGAGATATACTTCGGCTTAGGC
 CTAGTTCGAAGGATCAAAACTACCGCTCAAAATCGGTGCCGTTATACCCGTCC
 GGTGATCC 1-5: TTGCTAGC AACCGCGGCCACCCCTAACCTACCCAGTACCCACAT-
 CTCCTCTGGCATGTTGGG AAACCTGGCTAT ATTGACTT 1-6: TTGCTAGT TACAAG-
 AAAGCATTAGACCACAAATATCACCGCATTAAATCAATGGCTAA AATCGCTGAT-
 TCATCGGTTAACAGACAGCGCTAGGTTACCGTAGCAGCACTCGATCTAGTC
 TCCTAGGA

2-3: GTTGCAGA AAACTTGAAGAACGCTAACGAGTCGGACCTTGGCACTG-
 CTGGCGAGATGCCGGACAAAATATGCATGTGCCCTACTTAAGGTAAATCCGAT-
 TAGAGC TAAATCAT 2-5: GTTGCAGA CCACAGGAGATATGATATATCCTCGGCC-
 ACTGAGCCCTAGGCAACCCTGGTCTAAT ATTGACTT 2-6: GTTGCAGA GTAACCG-
 CGAGCCCTAACGTTACTCTTGACGGGAGTCACGAACGTACG TCCTAGGA 2-12:
 GTTGCAGA GGCACTTGA GATGTCGGTG AAACCTGCCT CCGATCTCAA CACTAGT-
 GTG GAGTCTTCAC ATCACAATGG AACTCATAACG GATAAGTCAG TCTGCCATGC
 CACAGGTAAA ATATGAGGCG AGGGCAG CTC TGGGCCTTG TATCAGCCAG GTTC-
 TATTGT TTCAGGGCCC TGGTTAGGCT GCAATCAAAT ACCCATAGTA TTTCTGAGCA
 CAATTCCAAA TGACGCCAA ACAGACGTGG ATAAATTGGT ATGAAGTCTA TGAC-
 CGTTAA ACGGAGTTAC TTC GCAAATCG

3-6: TGACAGTA GTTTGGACAA TGCGAAGTC GAGGTATTCA GTAGCAGG-
 G TACCGACAGA ATATGTCAG AAGCTT TGTT GGTCTAAAAA GTCACGGCCA A-
 TTTGCTAGC CGGGCTGTT ATACGCTCAA GATGGGCACC AGATGATATC CTCA-
 TCATTG GCCCGCAAGT GACTCTTCGA TTATACT TCCTAGGA 3-11: TGACAGTA G-
 TCCAGGAAC CCCTTTTTT TTTGAAAGTT AACGACCGTT CTACCCGACG TTAGG-
 GCTGA CGCCTCCGT ATACCAACAG CAAGATTACG TGATGACGGGA CCAAATA-
 TA G CTCGTTGCCA CAAGGGTACA ACTAGGCTTC GACAATGTAG AGGAACGCA-
 A CGGGTCTGTG TCCTTCACAA ACCTACCTT GGGAGGGTAG TGCTTGTGTT TAT-
 ATCTTC TATGAGCCAG AGACTCGATG GCCTGATAAG GCTGCCATCG GCTAAT-
 TGGG TACTTCGGCG CTCCCGAATA GCCAATTGT GTCAGCGAGA TACGGCCCT-
 T GTGTTCAGCA TTGCCCTGTG ACAATTCTT GTACGACCGA CGCCCATCTA TGG-
 GTGCCGG GGTGGTTAA AGAATTCTAG GTGCTACCTG CGACTCACCA CCGCTG-
 TTAT GACGGGGATA GTCGATACTA AAACATACCAT CATATTCTAG CTCCGGAAAG-
 A GCCGGCCAGT ATAGGTATTG AAGTCCTC 3-12: TGACAGTA CTCAACTACG TTT-
 AATTAT GCGAAGCGAG TGATTAACCG AAGCGAAGGG TAGTACGGCT TCTACC-
 GCCA GAACACGTAG ACAGCGTGG G AAAGCGATAG TTGACAGCTA AGTCGGTT-
 GA GGCTGCACCC TGATGAAC C GTAATACATA GCTCAAGCTT CCATTGGTGG A-
 TTCTTACC GTAGTACGAC TCACCCACAG GAGTCGACAC TGTGGCTTC ACTTC-
 ATCTA TCCCAGGGCT TTCCCCTTAG CAAGGCATTA TTAGACAGGG TATTGCCGG-
 A CTAGCTAGCC TTAGCTATAA CAGGAAATAA TGCTGATTGG ACTGAACAGC G-
 ATGTGATTG CTCCAATAGC ATATATCCAT CGCCGAGTAT TCTTCGGTTG TTGCGC-
 CATA CTTCTCTAGGA ATACCGCCGA TTATGAATTG GCCCCAGGGAGTCCGTT-
 C CGACCCATGA ATGATTCAC CGGTGCGTGC GCTCGGTTG TCATCTGTG GCG-
 AAAGGTC GCAAATCG 3-13: TGACAGTA AAGAAAGCTGG CCACATTGTC CACCTA-
 GCAT TGATTGGCGA TGCGGTGAG GGAAATTGCC GCGGTACCAA GGCAGAACT-
 T GCGGAGTCCG GTGAAGTATG GAACCGGGAA CACGAGCCGT AGCTAGTTG AT-
 GCTTAAA AGCTTGAGAA ACAATACAAC CTGCCTGAAC CACGCTGCAG CAAGA-
 CGCGA GGCGCATTAA ATGGCGAGGT AACCAAGTAT GGTCCCCACC AGCTAACG-
 AG ATTAGCCGAC AGAGCATTTC TTGATGCAA CGATTAACGC GCTGGTCGCA A-
 TGGTCCCAA GGGCGAGGGAGCCGGTTA TCAATGAAGG CCATTATGGG TAAT-
 CCCATA GACCAGGCTT GGTCAAGAAC CTATCCTAGT AAGTCGGACG TTGGACGT-
 TG CAAGAGTGC AAACATGTCC GCATAGCGTT TGTAAGAACT ATCGAGGAAC T-
 ACCTTGTC GATTGTCTAG AACATGCCA TGAAGAGAGC ACAGACATGC CAAC-
 CCCAAA TTATGTAGGA AGGGTACGGT TCTAAGGTAT AGTTTGAGG TTGTTCTTT-
 T CGTCTACAGA AAAGATCCCC TCCGGGAGTA TAGTGCAAAA TCCATGGAAG A-
 GCCCACAAA AGTCAGAAAT AGACTAATAT TCCATGTTAT CAATCAAGTT AATG-
 CTTCAA CAAACCCTGA GCGTGCACGA AGAAAAGTAG AGATTAA ACATATGT

4-5: GTAAATGA GAGAGGATTCTGAGCTGTT CAGGCAGCGA GGCATCTAG-G TGGTGCAGCGA AAGAGATCTT CTTGTGCTCC ATCCCGCCCC TACCACTTAG ACC-AAGACGA ATTGACTT 4-7: GTAAATGA TAGTGTGCTA GACATCTGGC CGCCGGCG-CC AGCTCGTGGA TGCTCATAAG GCTTAACCTG ACGTCTTAGG AAGAAACTCA A-GCCCAATCG CATTCTTCTT CACTGCAACA CGCGGTTGTT ACCGGCCTT GAGTA-TCTTT TACTTCGCTT TAGGCCACC TGCAGTAACC ATCGCTGTCA GAAT TAGCT-G AACTTTG CTACACGA 4-14: GTAAATGA TGCGGGTGTCTAAGAAATT GCATG-AATAA GTGCTGACTA AGTTATACCG AAAAAGTCTT GCGGGACGCA GTTGCTG-TC TCGTATCTCT GAGAGCTCGT GTCCGTTCGG CGGGGACGAT AGGATGCCCG CG-ACAATGAA CATGATCTTC GACCCACAAC GTACAAGACCG CGAGCTAACATGCT-CTAGT CATCCACGTA TTATTATTAC TACCACTAGC CAACGTGCAT CCCGCCTG-T AGTACGCAGG CTGTTTCACG ATGGCGATT CCAACGGAAA GTTCTAACCTT GACAGCGTAC CACTAAAGGA CGGCCAGGCA TGGGCATCGC AGTGGTGGAGT ATTCA-CTCCG TGCACAACCG GACCTGACGT ACCTGAGGAC GAGGTACCCG GAACTAGA-CT CTATCCTCCG CGGGCACGAA CTATTGACTT CAAACGTACG GACACCTATC TT-AATAGAGG ATTTGACCGA TACTCGCTAG CTGCCGGTT CCTCGCGGC GTTGGT-AT

5-6: CCGGGTAA CGCGAATCGC GGGTAGTTCG TATCGTTGGA TGACTGCAC-T GCACATTGTC TCCTAGGA 5-7: CCGGG TAA ATGTCATTC GGTAAGGACG TGG-TGACATT GATGCATCTG CTCAGTGTGC CATATTCTG CTCACCTCTT GCGCTATC-CTT TCGAGGGGTG TCACACACCT CTCAAG CTACACGA 5-8: CCGGGTAA GTGCC-CAGC TATCTGCCGC AGACAGGCAT ATAA CGGAAG GTATGGTGCCTGTCAGAG-CT TTGCGCAAT ACGCATGCA TTGGGAAT 5-9: CCGGGTAA GGCTCAGGCA CAA-CTGAAGA TTGACACACG TATAGCCAGC CGACCAACTA ATCACCACCG CGTGGG-CTGT CTCGATTTG CCCACTATT GCGGGTAGA ACGCGT TACGGGTT 5-14: CCG-GGTAA TTTCTAGCGG GCCGCCATCT TACTGAGTAG ACAAGGAGGG CATTCTGA-AC TTAACTTCAC TATCGAAGCG AGGCCCGCGG GCTGACAGCG ACAGTACACA T-AAACGTAGT ACGTATATT CACCACAAATG CCTTGCCACC GTCACCGCGA GTCTA-GAC GTTGGTAT

6-8: GGGTAGGA TGTACCTCG CGAGGGCGCC AATTGCTAACG TACAATAAC-C CCACGTCTAG GCTATACACG ACGATCTAGC GTTCCAGTCG ACTCGCCCAG CT-CACGATAA TGTGTACGCA ACAACGCCGC TGTCGAGCTC CAT TTGGGAAT 6-9: G-GGTAGGA GCGCAAGACT TGATGAGTGA TCGTCCGTT TGCCCGTGA GATGC-TGAAG CTCATACAGT ATGGCCCGCG ACAATCACTA TCGGGTGAAG TGTGGCCC-C TACGGGTT 6-10: GGGTAGGA GTGGCCGGCT CCGCCAATCA GTCCACACGT TGA-TGCCTCG GGACGACTCT CGCGTAGAAG AGTATGACT GGGACCCAAA AAGCTT-GAGA GGATTTCGA ATAAATCA 6-12: GGGTAGGA AAGGAGAGCT CGGGGGGGAA-T GTTATGAAG AAATGCAGTC AATACACTCT AACATCCTAC CCGAACAAATC AT-GATTAAC GTAGTAGTT ATCCTATAAT TCGCTCCGCC CTTGCCCCAG AAGCGA-TCAC CACCCACTAC AGTATACTGG GCTAACATAC GACGATTAAG TACCTTGTG-C TAACCTCAAT TTCGCTACGC TCGTTGGCT TATACCTACT AAAGCGAGGT CTA-CCACCTA CCCTAAAAAA GCAAATCG

7-14: GCATCGGA AGTAAATCAC GCCTCATCT GAAGGCGAGT CGACCTAC-TT TCAGGTGCT TTAGAATGCT ATATGAGATG ACGGAGGCAT TGGTTAAGTA C-AACCTGACC AGCAACAACG AGTGGAACCG AGAGCGCGTA CAACTATATG TCA-CCTCATC TAAAAGAGCT TACCGAGTTAC TAATGTACGG TGGGGAAACG GCTGA-GTCGC CCCTCCGGTT GCTCCCCGGT AGAAGCCCGA GACGAGCACC GTCCACA-GGG GTTGGTAT 7-16: GCATCGGA TGAGCCAGGG ACTGAAACGG TACCACCCAG

TCGTGTGTT TGGTATCTTG GGGCCGCCCT TATCGCAGCG ATGTCTCCAA GAAG-GTCGTC AAAAGCAGAT GGAAG TTGAT GCATTCTCTG GCCGTAATGT ATGTAGG-ATA ACCCCTTAGT TCCAAACTGC AATCACCCAT TGATAGTTCC ACATGTCCTG G-AATTAAAAA GGACCCCTTAC ATCCTTAGCT TATGAGGTGC AGAATCCTTC TCTGG-CGGCT ATGTAGTAAG GTCACGAACG TAGGTAGAGA CAAATCCGAG TTGATTG-CC ACACGTTGGC CGAGGCTCCG AGAAGGGAC TTCAGACTGT TCTGGTTACC G-CTG ATCCATAC

8-9: GCACGTAG GAATGAGCTT AAACTAATCA GTAAGGACCA GTCGGGGA-AA CTCTAGCCA TTGACCACCA GTTATTCTCG TCGGGTCGAG AGGCTCACAG A-TAATTAGA TACCGAGAAG CGAGTCTTAG GGGAA TACGGGTT 8-14: GTATAT C-C CGGTACGATC AGAGACCCCTC ACGCAGCAAC ATATAACAAT GCGGAAAATC T-CGTGCTCA AATGCGTACT ATTCAATATCT GTTGAACACT ACTAGGCCGA ACGC-AATGTT GTCAGGCCAA GCCACCGGCT ATAGGGATT ATTTCACCCCT CTAGGC-AG G ATTAATAGG GGCAACCAG TTGCTGGCC TGGGTATATG TAATGCCTA-A CCATAAACAG GAGGGAACCT GGGTCAGCC ACACCATGCA GTCCACAGGG 8-16: GTATATCC GGAGAAGCGT GGCACCCCTG GGGGTAACTC TACATTCAACC CAG-GAGG GCC TCTCGCTTT CAACTGGCA ATCTAACATG TGTCCACCGT GTACCG-GTAG CGCTACTGCC CGCTTCAAGA TGTAAAGCGAG TACGCGTCAT CTGTTAGCG-A TGAGCGATT TAGCGGGACG AGAACAAATAG TAGAGTCAAA CGGCTTGGAG C-TAGAC AGCT TGTTCTAAA CAACACTTAT CATTGCAACG GTGGGGTAA GATA-TTCTT AATGGACTCG CTAATATATG GGCCTGTTA TGAGGACGCG GAACGTA-AT CCACCAAGGT CCGCCGAGTC ACATTCTGC CTATCGTAAG CACT ATCCATA-C 8-17: GTATATCC ACCTCGATGG ATTCTGGTC TACCCAGGCA ATCGTTGATT CC-CCAATGGC AACTCCAGTA GGGTGTACC AATTGACAC CTCGCAGATC AACT-GGCAC GGTGAGGCTC TCCGGCCGAT GATGGCAAAT CGGGTAATT AGATGGGT-AC AATTGTTGG AGAAAATCGT GGCGCTTGC TTGACACCT CGGGATCGGC A-AATCCAGTA GCGTGTATGC TTCGCATCCA GCGATGTAAC CATCCTGAT TAGAC-GCAAA AGAAGATCAT GGACCGGTGC GGCGGGATA CAATCGGTGT CTGGTGT-CA CCGACTGGTC CACCATCCCC CGGTGAGGGT ACGCGGTGA TGAA TCTTAGC

9-10: GTATATCC TGGGAGCTGA AACGAAGTCG ACACACTCAG CTCACCGC-TA AGCTAGAGGG GGAAGTACCT TTGTATGATT CGGCTGTCA GACTATCGAC C-AGGGCTAGG ATACTGACGG TGCAGATCAG GTAGC ATAAATCA 9-17: GTATATC-C GTACACAGCA GCGAGTCACA GACTCACCCC CACCCCAAAA GCATCGTGAT C-TGTTGCCGA CTTGCCTTTA CGCGTAAAG TTTTACTTA GTATCTTAAT CTCATG-TAGC GTTATAGATA GACAATTATC AGATCAGTAC TACAACGGCC CAGTTCT TG-C AGCGGATTGC CTATTAAATT CGGACTTGTG AGATTGATG ACCAAGTTAT AA-AGGGTCG TATTGCAAT TCGCTGGTCT ACACCAAGCG TCTTAGC

10-12: TCCTAGGA AATGTCCGTT CGGCAGGTTC TACGTTAAC AGGTACGT-CG GGTGATTCA CTATCCGCGC TCACCCCTAGC CATATCGATC TCACATCGAT C-CGGTTAACT CGTTGCCGT GCGGGAGGCT GCCGTTCCGG TAGGGTAGCA CTTT-G AGGAA GAAGCAGACC GAACACTGAG TGTGAGACCC CATAGCTGAA TTAAA-GCGAG CTTAGGGTAG GTGACAGCTG CGG GCCAATT TGGCAGCGTC GGACTGA-ATG ACATGGAACC GGCCTGAATC TGCCTGATTA TGAAGCTATC GCTTCTGGTT-CTCAAAACA AACTCTCAGA TGTGTTGCAC CCACAGGTGCG TCCTTGGTCC CGAAT-CCAGC CTCATTCTTG GGATCGACTT AGAGGTCCAA AGCTT GCAAATCG 10-17: T-CCTAGGA CCCAACTTAC ATAGGGATCA GACTAACTTC ACCTGCTACT CTAGAG-CTGT CAGGTACGCC CACTCACGTG AAGCACACCA GTTACTCGT CTGAACGTT

GACGAGATAT TCGCCGGTAT AGATCGCGCC CTTTGATTTC TGTGCTCCCA AAAA-GGATT AGGTGTCAGA ACTGGTACTC CCGTAACGTT GGCTTTAAG AGAGGTG-CGG CTGTCGGGA CCCTTCGTCG TGTCGCTCCA GGCTTCTGG GTCGACATTCA-AAA-GGAGAAC AAACAAAA CG TGGAAAAATG AAATCGCATG ACTTGCATGT TAC-TTATTG CGGGACGAGA TTTAAGGAA ATTGGTACCT CTAG TCTTAGC 10-18: T-CCTAGGA GATTACACCA CACGTACACA TAGCTCCCA GATCCCTCAG GTCAGC-TAAA TGACGCCCCC ACAAGACGGC TACGCTTACC ACCAATCGTA TCCCGTGC-A GTCCAGATGC GGCCCACACA GGCATCCAGA AACTAATTG ACGGTTCTAC C-ACGACTGTC GAAAATTGCG ACGCCAATT AGAACAAATAC CGTAGACCGC GCCT-TCTGT CTAATTCTGT TTCCTCCGGA AATTCCATAC CGTTGGATAC TCCGCCGT-A AAGGGGTATG CTTCCGAACC AACAAAGGTA GATAAACGCG ACACCACGGT C-TATTATACA TCCCCTGGCA ACCCCGGGCA CATCGCTCGT TTGA CTCGCTGA

11-12: CACGTGAT TAATATACAA AGACAAACACT GGCATTCCAA GTTCAGT-TCG GTCAACAATC CTAGATGGGG CGAG TTCATG ACGTGGAAGC CCATCAACA-G CTGCACTTCG GGCAAATATG GTGCCAAGG CTAGTCGCAC ATTACGACGA G-AG AGGCACT GCCCTAGACC CAGGCACAGG CGAAGGAAAG CGCTCTGCCG GC-GACCATA ATAATGACG GCTTAAGGTG GAATGACATA AAGCCGTCCC CCCA-ATCGGC TTATATGCAA GAGTACTGGC ATCAATCCTT TAGAGGGCTG TGGGACT-GGC CGGTAAAATG GCATTTAGCC GAACTGACTA CCGCACCAAG AATGACTAG-T TTICAGTAAT CGTTTCTTT ATCACAGGAC GTTCCTCTGG GCAGATTGT CAC-GGTCCGC GACGCGCACT TCTGGTATAA TACTATGATA TTGAAACAA CATGCC-AGCC TCTACCGCGCA CTTGGGAAA CAAGGCTTAG GTCCAGATGA TTTCTCTCC-A CGAGCACACC TGCCGTAAAT GAGAGATGCG TTTAGGAGTC GCACCTATAG G-TCCCTGCC GCACGAGAGT GTTATTCTT CGCTCTGATT TCGAAATATT TCACA-AATGT TCTAAGCTGC TTATGCTCG TGATTGACC CCCGTTGGTT AGCAGAAC-T GCAGTCAATG AAGCTATTCC TCCATTCAAC TGCAAAGTAA ACAGCGGGGT GA-TTCAGGGA TGCAAGTCGA TGGCCTAGTA ATGTGAGGTT TGATAAGTCA ATACTT-AT CG GAGTATCATT ATTGCTTTT GACCCCTTGG CGGGACAAGA ATCTACAAC-G ACGAACCGCA AGATAACGTT TAACTTACAT TGTCACGTG CGCGGCCGGC AA-AATATACG CGAGAAAGTC AGCTGTGTTG GGCAACCGGC CAAGGACTTG GACG-CACCTC ATCGCAGCTT CGCACGTAAT TGGAGCGACT GTCAACTCAA TAGGTGC-CCG TAACTCTGC GCAAATCG 11-13: CACGTGAT TGTATCTCGC ACACCCTTA A-CGTCTACCT TAGAACCCCC TGCTTGAT CATTGTCTG AATGGCGGGC GCGAT-TGGGA GTTCGTGTTA TATAATTGCA GTCTGGCAAT AGAGATTG CGCAAGAA-GG TACAGGGGCC CGACGCCCGC ACGCCCTCAC ATACCTTCTC TGTAAATAGCA A-CTGTCGCTC CGGAGCTCGG GATTGTACCC ACATGGTTGT GGTCTCTATG GCGGG-GAAGA GAACATGACC TTTGACCTGT GTCCCGTAGC ATCTTGGCGT TCCTACTTG-A GTGGATATTG ACCGAATTAA GACATACAGG TAATCTCGGC CGTTGGAGC GC-AATCCGCC GCTTACATTA TCAAAAATGA ATGCCCGTA CCACATATTG GTCCC-AGGGT CTTCTAGACC CGTCCTGTG TATAACCGAA CCTCAGAAC AAGACCGC-CT ACTACTGACA ATCCTCGGCC CAGAGCATCC CTAAAAGTGG TATTCTAAGG A-CGCCCGCT TTACGATTAG CGCTCATCGT TGTATTAGT TTAAAATACT CTTAA-GCGGT TGATGCGTAG GCCGCAATGG TTTCGCAAAG GCGTGGCACT GAGCTGC-TGA ATAGTTGGGA GAGGCAA GT GGAATTCTG TTCCAAGCGC CAGGAGACC-G CCAATGGAGC TTTCATTCCC AATAGGGACT TGATAATAGT CGGCGGCCATA G-AACGCTGGA TACAGCCATA TCGCCTGGCC TCCAGCCAGA AATCGTGCAGT-AGTAG GCAACTCACG TTCGAAGAAT TGGACCTCTA GTGTGGCTGA AAGCTCG-GCG CCACCAACAT GCAATTACTC TGCCCATATG CAGTCCGCCA TTGGCCCTCA

ATATAACCAA GACAAGGGGA CGGAGATGAA CTCCACTGGT CGGCCACGTT CGA-GGGGGGG CATTGCCGTG TGGCATCCGA TGGGTTCGCT TCTACCCTCG AGTGTTC-ACC CAACGCTGTA CTGGGGTTTC ACATATGT

12-13: GTGGAATC AGGACCGCAC GGTCAATTGT TATTGTAACA CCTTGACA-
GG CGGCAACGGC TCGCCGGCAA GATACCGCAA GAGGAAATGG CGATTGAGG-
A TCTCAATTCA CTGGCCCCGA CTCCAGTTAC TGCCTCGATT TAGCTATGTT GCG-
AACTACA CGCGCTAGAG CAGCGTAGAT ACAGCTGACT CCCCTAAGCT GTTCCC-
GCGA AGCGTTATAG CGAATTATA GC TCGGGCAT GGTGGATGGA CACCGCCCG-
G CTGACCGGTG CCAGGCCATG CAAGTCAGGC TTCTTGTTC AGAACTAAAT AG-
CATCACAC CAAAGCTAAA CGTCGGTGG GGTGGTCGA TGTGCCTAGT CCTTC-
ATGAA TGGATTCAAG CTATTCAAG AGTTCCCGC CACGAGGTCG CGATGTAT-
AG GGGTTAGCAC CCATCACACG GAATCGGTGT TCTGACTACT GGAACGGTCG C-
CAGATATTG TTGTACCGT TGATAACGCT ACCGTAATCA CCATTACCCG CACCA-
CTACG ACAGGGTTG GTTTGCTCC CAGGTTGTC CGGGCGTCT GCTAATAGC-
G GCCTGGCAGT TGTCTAGTTA CTAGTAGCAT GGATACGTT GGAGTTACGC GG-
CGTTAACG TTTATTGGTC GTGGGCTCAA CATCGGTACC ACTGCGGCC CACCAG-
GCGC CGTATAGCCG TGGGGGATG AGGATTCTGC ACTACTGTAG TCCGACAGG-
C AGTGCAGG AATAGGATGA AGGGCGACCC GAGTTCGTCA AAATCCGCAC A-
CTATCAACT GTCCCCCGTC CCTGCTACG GTGGAGGGGG TAATTCATCC CTGAG-
AGCAT AAGCAGTCAG TCAGGC TCGG GCCGGATATC GCATCGATGT ATTCCGCA-
TT CTTGCTTCGC GGGGTGTCG AAACATCTG CGCCCCCAGC GACAGAACAT TA-
TTCAAGTG GAAGCAGGCT CCAACCAGAA AGCAAACGGA ACGAAGGGCC AGCT-
ATAGG ACATATGT 12-15: GTGGAATC ATACTAAGCA CCCTAGTTA AAGAGTAC-
CG CCGAAAACAT TGTGAGGAT ACACTGCGAA TGCTGATTGA GACGGGAACC T-
GCGACATAC GGAACCAAGG GATCTCTATT AAAGCCGCTC GCTACTGACT AGAT-
CCCGTA GATGTTACGC GACCTTAAC CGCTGATAGG GGATTGCCG TGGCGTT-
GT ACCACTGCGC CGACTTCTCT GCCGGCGCCG CCTCGGGTAA GTAACGGTTA G-
TAAGTCAG TTGGTTGGG CCCATTGAG TGAGGTCCT GCATGGTTG ACTCT-
CATAT GCCTGGGAG GACAAATTGA TCCCAGCAGA ACCAGATTG TAGGATGA-
TA GTTGATAAAA CACGTAAACA CTCCTCTATA ACTGCGCTGA GGGTGTCAAC A-
GACTGTAT TAGGACCGGG GGCACACATC CGTCATATG GCGTGGACCT AGCCC-
GAGGT TTAGGTTAG TTACCCAATC AAACGCCTCA CATCTGATGA TACTTACGC-
A TTGCCTGACG AGTTGCGAGT TCTAGGCTAA AATAAAGATG TCGACAGTCG GT-
GGGATAAC ACTTTGGTGT GAGCGGATAT ATCAGAATGG CGGGACCATC TGGGA-
TAAGC CCTCATGTTG GGACCTGGC TGTCGAGGCC ACCTGCATGA ACTGCAAG-
TT TGACTACGTA GGCATTGCC GCCCCTTGA ACCAAACCGG CACTTACTAC TT-
GCGAAAAAA AACGAGTCC GAACCCAGCA TCGATCCGT GCTACGGTCG AGTAC-
TCAAT AATGGCGCTA AAATAGGTAC CGGCATCGTA GCCCCCTCGG TTAGCAAT-
TC TGCAGTAATT ATAATATGGC AATTGAGGTT GAACACTTGC TATGCGGACC TT-
TACGAAGT TATGGCAGCG TGCGGGTCAG CTATTTGGT AATCGCCCGC TGGTGT-
CTGA CCGCTACACA AACCGAATAC CGCACAAATAC ACTGCCACAG CACTACTAT-
C AGACTAAGA AGGGCAG

13-15: GAAATACA CCGCGTTCGC CTGTGCGTAA GTCTAACAAAG GCCCCGC-
GAA CAAACTATAT CCAGAAGATA GGTTCCCAGC GTCTGCTAAC GAATGTGTT-
T TATTGACATA CCAGGCAAGA GCTACTCCCC CTCAGGCCA TTGGGGTAAC G-
ATGATTGCG CATTAGGCAC CCGATGACTA TCGTAGACGC ACACCGTCGT TCGT-
CTACTA GGACAGTTT AAGATGCTCC TTGGTACCAA TTGTGCAATG GGCTGAA-
TCT ACACCTCCTG ATCATGGACG CAAGTTATTC CAATTAGACG ATTGGTCGAA

ATCCTTCCAT TGACGACCTA AAGTGAATGC GGGTACCTGT ACGAACCTGT CCC-
CCAGATC AATAGGACGA ACTCCGCGAG ATACCTAAC CAAGTTGTCC GCCGC-
GGCTC TTGGAACCGG GTTTAGGTG CGCCAATACG TTCCCGACA GCGAGTTCT-
C TTAGAGCAAA GACAACAATG CGTTGTGCAT ACCGAGCCAA GACTCCGAGG A-
CCACCGTG CGTCAAAGGT CAAGATAGAT GCGACGTTT TGACGTCTAC CAGA-
ATCGT GAAAGACACA TCCCCTGGCT GTTCGCCCTA GTTGATGCC TCCTGCAT-
CA TGGGATAATC TCTTCAGCA CCACGCCGCC GCTACACTAA GCGCCTCAC A G-
CCTGAATAC CCGCTTTAA GCTTATATG GTAAAATTAG TAGGCTTAG ACTGC-
TTATT GCGCGCCGCG CCTCGCTGAA CCACCGACAA TTACTGCCG TGAGATCAT-
T TGTGTTACTT GAACCCAAAC ACAGAGAGGG ATGCCAATGT ACCTATAAGC AG-
TGATCATT ACCCATAGCA GGTAAACGAGG ATACTGGACT TCCCAATTCA CACCCT-
TCAA GCGTTGGCGT GGGGGATGCG GCCTATGCAG GGCTGAAGTC TGGAACCCA-
A GAGAATTCCA AAGGGTATTG AATTGAGCGG TCGCGATCAA TAGTTGTTA TT-
CAGGGCG AGGGGCAG

14-16: CGGCGGCA ACCTACGCAT CAATCCACCC AGTGCCTATT CGTACATG-
AC AGTTATCGTT TAAGATTCCG ATAAGTACCC ACATAGGGGG GGTTGCCGA T-
ACACGCCGG TTGCTATCGT CTAACATAAC TGATAATAATT GACCCCTCAG TGAGC-
CGTAT GCACCGACTTC GAACATTTT CCGTGTTCAG GTTTGTCAC GACACGAAA-
A CCTGCAGAAC CACATGGTGG CAGAGCTTGC ATACACCCCT TGATGGAAA G-
GACTCTAGC TCTTATCTT GCTACGGCAC GATTCGGTT TACCGTAGAT TTAGA-
CCGCT TGACCTTGAG TTCCTTGATT TAGCAGGTGA ATCCATGCGA AACCCATGC-
A CGATTCCGTT CGTATCCCTA TATTGTGTT ATGACACAAG TTACTCAGTT GTCG-
GGGATA CCGGATATCG AAGACACGAA AATCGTAGGC CTTCGTTTC TGCTGACT-
TG TGAGCATGGC AGACCACAGT GAGGAGTGCC ATCCATAC

15-18: TATGGAAC TGACCTAGCA CTTCGTTGTC GTTAGCACGA GGGCTGGT-
CC CCTATTTGA GCGACCGTGC GGCATGCGAA GAGACCTCAA GGGTAGTTC G-
AAACGGCTT TAAGTTGAGT CTGGATTGT TCCTCGGTA TGTGACCGGG TTTAT-
CATCG GTTTACCGG ATTAGAGTTA ATGAAGAGCC AGTCAAACC TGCTTCAC-
G AAAGTGTGGG AGGGCGCTAG AGAATAAAAG CCCTGAGTAT AGCCTCGCG C-
GAAATTAGT TGCACCACTC GTCTACGCCA TTCAACAGA TTAGGATCGT ATCCT-
TTAAT GGGGGATGGA GGTTCATTC ACCGTTTGCA AAATGGCGGT CCGG CTCGC-
TGA

16-21 TGAGCCGC AACGACATAG CAGAAATTAA CCCCCCTAAAGA CGGCAA-
AACG AGAGAGCATT TTTAAGTAC GAGCACGTAT TGTTATACTG TTAACGACT-
A AAGGCAGAGC ATCGTGCTCC ACTAGTCATA GGGTGCCAGT GTTCACACCA C-
TC GTGTGCA TTAAACGTCG CGAGATAATC CAGACTAGCA GTGTAAGCAG AGC-
CACGCTA GGGGCGGGTT CGCGGTGAA GCGCTACCAAG GTCGACCTGT CTACG-
AAAAT AAGTAGAATG GTGTGTCGTA TAAGACCATT TTGTAGAACG CCGGGAA-
CGA AGGACACGCA CAGAGGGGT A TGCTACGTTA ACTGCATTGG TATCGGAAC-
C TGAG TTACGTGT 16-22: TGAGCCGC TAGTAGAACAA TGCCCTAACG TATGCATG-
CG CGCCAATGCG GCGACGGAAA CCACGTCGCC AGCTCAATAG AGAAAACCG-
T CCTCCGCC ATCTGACA

17-22: CGGCGTTA GAAGAATAGG GCTTATTCGC GGAGGGACGC TGGGATA-
CCG TTGCGGGTAG CGGTTACGCC TACA ACAACA CATCTTGAAG TGGGATGGC-
T TGATGAGCGC TGGTGAATT GTTTAGTA ATCTGACA 17-23: CGGCGTTA GGTG-
GAATGA GGTTTTGA GTAAGTCGTA CAGTGCCCTA GCGTCGGCG TACGCCCG-
TT CAGCCACCAA CCAACTTAC GGACCTCAA ACTAGCAGCG TGGTCTTCAC G-
TT AGATGGCG

18-23: AGGACCCA GGTGGCGATG ATCCCCCTAT CCGCAGAACT CTCGCGCTTA AGACAGTCGA TTGGGCAGCA CAAA GTCAAGG TCGACTAAAT CCGCGGCCG-C ACCAACATAA GGAATTAAAA ATTGCATGG TTCGGTATTA GGAATACCTA CATGATGAGT AA AGATGGCG

19-21: ACGCCTCA AAATCTAACG CCATTGGGG TAATCGAAGC CGAAAGACA TCAGCACCAA TCGGCAAAAAA GTTG ATCCCT CAAACATACG AGCTGTGGA-A ACGACTAACGG CGGTGCCCG ATGGCATACT CCTCTTAGA AGGTAAACGT T-GAGCTAGCC TCGTTGGGAT TTGCCGTTT TAAAAGGGGA GACTGTGTAC CAGTT-TAACT CGTTGGTCGT AGTACTCCAT CCTCGGGCA TCTCCATGT GTGAGAATA-T GGAGGAATGA GACTACTAGG TCACTTTGGG AAGCAGCACC CTACAGAAGA G-TCGATCGGC TCTGGACATA TTGTTGTGAA TAT TTACGTGT 19-24 ACGCCTCA GT-CGACACTG CGTCTGAACG GGTCTGGCC ACCACCGAAT CATTATAAGA AGCAC-ACTCC ACCTCGAAG TATCGATGCC CACCAGCTAG AACGACTCGC ATCAGTGA-GT GTAGGGCCGC ACCGACCGAG TGGAAGTTCA GGCAGCTTG CACCAAATCC C-CTCCCTGA CTGCGGCAGT CGAG CGCGTC ACCAGCTACC CATTATATA AGCA-GGGAGA CGATTATATG CGGATTCTCA TAATCGCACT TCCCCCGTAC ACCCTCGG-AG ATAGGATCGG AGACTTGGGA AGCGTAGTCG TCTACAATAG ACGTGTACA T-GGTCTCGG TTGAAGAGGA TA AACAAATCC CTATACCTTA CTTGGCCAAA TGCG-TTCATT GCGGGCCAAT TCGCGACCGA GCAAATAAGT AGTGAATGA TCTCGTC-CCG GTATTATCCC AGCCTAGCTT CCGTCTAGAA CGATTGTTA GGTTCAACGT T-AGTCGCTA GTAGGTTATA TTACTTG 19-25: ACGCCTCA TCGGCCCTT GCGTACCGAA ACCTCCCGTT TTAATCTCGC AGTCGGGCC GCGAAGGACA TCCTCCCG-G GGGTTCTATA GAACCTCATC AAGAACCGGC TCTGTGTAAG TCTCCTGGT CTGGCGACG AGTTACGGGT TGGCTTATGT CCGGTATACA AAAGTCTCCG ATGATGA-GTC TTCCCTGGT AGTAACTCCA TTGGGGTCT ACCGTCGGTC ACAGCTATAT T-GGCATTAGT TACGATCTCA CATGTCGCGT TCGTACGTT AATATGGCGG ATCGA-AGTAT TGCACCTTAC CGCAGACAGC GCAGGCAGAA GGGACATTGC CAATATAC-GC CACAATAGGC AATA AGACGTAT

20-22: TGGGAGTC ACAACGCCCTC GAAAGTGGTA TTTCCAGTTG TTACGGAG-CT TCGGTAGTAA AAGGGTGGC CTGCTCAAGG TGACCCATGC GTTATACGG A-ACAAGGACC AATGGTTAAA CAATGGCCA GACGTGTATT TCGCGATTGT CCTA-GAAGAG GGAGCCA ATCTGACA 20-23: TGGGAGTC CTTACACCCC CTCTACTAC-T CGGCTGAACA GCTATTGGGG AGGTGCATGA AGGAATCCAC TTTGGATAAG TCCGGACCCCC CCGACAAGGC TGTTCCACG CTACATGCGC TGACGTTCTT ATAGCA-TTTC GGCTACTGAA GGA AGATGGCG 20-26: TGGGAGTC AGGTGCGCGG ACAGA-TGGAG ATTTGGCAC TGCGTACCT ACTGCAGCCC CGTGTGCCAG AACTATTT-A TTGCGCAATC AGATATTACA ATTTAACGGG GCGAAGGAGG GTTCTTCCC CCA-AAAAGTGG ACGAGATGAT TGAATTCTTA GGCTCCATCG GACGACACTC TTGAC-GCATA TGAACGCCCTG TAGAAGAGAC TCGCAACA

21-22: CTCAAATT CCAACGACAT TAGGGTAATA TCCAATACTC TCATCGT A-TCTGACA 21-24: CTCAAATT TTTGGTAA AG GCACCAACGG GACGTGAGAG TCCC-CGAAAT AGAAGCTTGT TGTGGTGGGA TCCCTGCCCG GTGAGTATTG TGGCGAA-CTC GGCCTATGTG CTAGTACGCT TGTATCCCAG AGGGTTCCCTT AACTGCCAC T-TGGAATGCC GAGCGAACCC CCTTATCTCC TGGATTGAGG AAACATCACA GGC-ATTGCGG GATAGGGTGA GATCAAGCGA TCCTTATGCG AGCCCA TTACTTG T 21-25: CTCAAATT AGGTACTCGG TTCTGACTTG GTAATAGCAT CGTACAATT AG-GTGGGAGA CAGTGGAGCT GTCTTTG CGC GAGAGGGTGT TGT AGACGTAT 21-26: CTCAAATT CAAGATCACG AAGTCCATGT TAAATCCCA TTCCCTCTAG TTGC

GCACTA AGCAAAAACA GGCAAAGGCT CAGCAAGACG AGATTGTCGC TCTG TC-GCAACA

22-23: CATTAGC TGGAATAAAT ATGTAT AGATGGCG 22-25: CATTAGC TGAG-GATCAT AGCGGCACCC AAGAAAGAAT AAGGAATGA AGACGTAT 22-26: CATTAGC TGGGCTGAAC TGTGCGTTCT CG TCGCAACA

23-26: GCAGGAAC GACGTGGATG GAGTGGCACA CTTCGTTGAT CAGTTATTG TCGCAACA

24-25: AATCACCG CGCGGCTATG GATTGTGGGT AGAAATAAGA AGTTGC-CACG GGCATCAACC AATACTCCTC AAGGACAATT TCGGGTCGTA GCTCTGGTA TAGCTGACGA ACTAACCGTG TATCCCACGT AAAAG AGACGTAT

25-26: GCGGGGCA ATATAAGGAT GTGGGTAGCA CCAAAGACCG CAACTTGC-CAA CGCCTCCCTC TCGCAACA

Next we present coding for Subgraph II.

VERTEX CODING:

1. GTTTTACCGTGGACAAT
2. CGCGTAATCACACCA
3. TAATGAGGACCTGGCC
4. ATTACCGAATAGCTAA
5. TCAAAACCGTGTGCGT
6. TGTGTTATG TGTTTACT
7. TTTGACCTCGTAGCTG
8. TACACGTGGATGAGG
9. GGTGTCGC CCAGCTT
10. TAAACGGTGCACCTCC
11. GGGAGCATAGCCATT

Finally the edge coding for Subgraph II.

EDGE CODING:

1-2: ACCTGTTAACGGAAACA TCATGGGTCC AGTCCCCACG GGACAGAACCGA-GGGACGGTATGAAGTTTC GCCAAGACGA GTCTGCATC AGCTGATCTT GTAAT-GTTGA CCTATGTTATGGGTCCATGG ATCACTACGA CCCCGAAGTG GA TTTGTT-TT TGTGGCGTGC CGGTGTA GCGCATTAA 1-3: ACCTGTTATGCGAACTT GGAATA-CGTC AATGTCTACG GGCATTGGTA ACTGTTAGCG CACCGTCCGC GACAGCAGC-A TACGGGTGG ATTACTCC 1-4: ACCTGTTAGGTTCGAAT ACCCTATGT AGTCA-TGAAT CGTTTATCTA TTGTACCCCTG AGTACGTATG GTAAAAC TAATGCCT 1-5: A-CCTGTTAGGTTCGTCA GAAGGTTTT GTTGCTGGT TCTGCCGCGT GCCCCGTTG TTATACACAC TGGTGGACGC ATGTGCGCAT GAATGTTCAAG CAGA AGTTTGCG

2-4: GCGCATTATGGATAACCGC GACTGTAGGA GCTGCAGGTT GACACGTTCT TACGGCTGCA CATATACTTA CCCTTT CTC TAGAAGGCTA GAAATAGCTCG-TAGCGCC GTCTCGTCAC GTAGTAACGA TGTCCGCATC TTAAACGCAT T TAAT-GCCT 2-5: GCGCATTACATACCGATA CGGTCAAATC TGGCAGATGG CCGGCTCA-TA TCCTGAGGCC GCTCGAGCAT CCCACGCACA ATTGTGACCG AACGGCGTTC C-CACGGCGAT AGTTTGCG 2-6: GCGCATTATGGACACCGC AATAGGAAGA GGGCA-CCTTA CTGCCGCTTG GACAAGTCGT GACTCGTAAT GTTCTAATAA CCTAATTTC-CTCATGAGTG AGTTGCTGTT ATAAAACCTTA CCGAATAGAA GCTATGTGCT GG-GAACGGCG TGAGTATAGA GACCACTTC CGGCGCATACTGCCGCCGCTATT-TATTAC ACCGAAGACA CGGGGCATTAGCCCGGAAT AACCGCGATGA CAGGATA-GTG TATGGGCTCTCACATTGCAT CGCAATGTTACATCGCGTG TCCCCTCAAG T-AGGGTGCAGACGCCATAAA CGGCTGACAC ATACCCGGTA CTAGATCACC GATT-CATACT AGCT ACAAAATAC 2-7: GCGCATTATCCGCAAGAACGTAAGTCG AGAT-TGAGCG TAGATAACAC CGCAACGATT CGGCTGTTAC AACCTAGGAC AGTTTCT-GGT TTCCAAGTAT TGTGCTCAGG TCATCGGATG GGACAGTCCTGGCGAGG

CACCTGGTTT CCTGCTGATC GACATAACGT GACTTGCTGA TGGGCTAAAAA TCAG-TACCAA GTACAAATGA TGCCAAAAAC GATTCACTA CTATGCCCTG AAGCATT-ACG CCCCCGAACG GGGTACAGAG ACCTCGGACA AGTGTACAC GTCCAGTCC-T GCGGGGACGT CAGTCTAGGG GCCCCTTCAG GCTTCGACCG TAATCAGTGA GA-TGAACCTCA ACTGAA TTG TATTCCCCGC TAGTCGCTAT GGCGAGACTA GTAGGT-TCCT CTGGCCATAC GTCGATACTA AGCATAGGAA TCAAACCCGC CGCCTACGC-T CGGTATGCAA GAACCCGCAG CTCTATGAAC AAGCTCAAAG CCTGAGCGGC A-AACTGGA

3-4: TGGACCGG GCAAATCTCA CCAACTTGAC TTACATTGCT GGG TAATG-CCT 3-10: TGGACCGG GATTGAAACA TTTAGTGACT AACGGCGTC GTCTTCATC-T AAGACTCAGG TGTGTGTAGC AAGTTACTGC GTTGAAACGG CGGAAACCAA CA-GCTGAACG TGTGTTTGA ATGGTCGCG ATATCACCC TCCACAGTGA TCCCTTT-TTG ACGAGTGGAC GGTACCC ATT TGCA

4-5: TATCGATT GGAAGTCGTG GGCAGGTTCT CCATAGGATC CTTAATGAG-G AGTTTGCA

5-6: GCACGCAC AGCCTTAAC CATCAAGTAA AGAACAGTGG CGTCATGG-CC GGCAGCTGGG TGTCCATGAG TTCGTACCGT GCTAGAAAAA AATCCTCGCG C-CATGCACGT ACAAAATAC 5-7: GCACGCAC TCTTATCAAT CAGAAGGGTT GGGCT-GTGGC TCAATTAATC AATAACGTT CTATCACCTG AGCCATTCTA AATGTGAAC-C AGGTCGCTAG CTAGATCCTG CCTCATGTAG CATTACACC CCTAACAGCA TA-GCCCGCTG TCGGGTGACG ACGGCCGTAAC CGCGTGCCA AGAACGGTAG GCT A-AACTGGA

6-7: ACAAAATGA CCTGGCGTC TTCGGCATAA CAGCGGGGAG CGCATACT-AT TGAGGGTACA ATCAGGATCC TTTCTGGTTT TCTCCGAGTC CCCTGGTGCC G-TGGTCCCAGC ACGGGGGTAG TGCCGGAATG TCGCATAGGA ACGTCAGTAT CTGGG ACCGC CCCTAGGTAG GTGAGTCCAC CGTGTATCGA GTGACCAACC AGGCC-CTTT TACTAAACGG GTGACATTGT TGCCAGATAC CTTCATGTT GGTCAGCG-C AAGCGCGTA ATTGCAAATG TTGCCAGATC AATCTAAGAT CACGTAGTGA G-TACG TCCAG CTCTAGAGCT ACCCTTCGGC TCTAGGAGCA CGTGGACGC AGGC-CGGAAT TACGCTATTA ACCTTGTAGT CCTCCAATTA CGAGCACCTC TCAAGGCT-GA GTCGTCAGGA GGTGCGCTT CCCTGGCTAC TGTGTGGCT CGCTGAGCTT G-TCTATTGCG CTGACTAAGC CCCCGTCTAA CTCAATATGG AAACCTGGA 6-8: ACAA-ATGA GGAGCCAACC CGTATCAGGT TGAGATGTGG TATACTTACG GCTGCCACTC-G TAGTTCTAA GTACTTCAGC TAACCTTGTC TCACGGAATC CACTTGCCG AAG-GAATATG CCTTGTAAAGA CTCCCTATA TTGAAGGGTA GAGAAGATTG TTCTCAG-GAT AACCCGGAGG TACAACGCAT CGGTGGCAA CCTAGGGTC GACGACTTA-C TACCGCTGGT ATCCCTGCG CGTGGATGG TAGCCCCCTG GCCGGCATAA GC-AGACTACG GCAAACGGTA ACCTTCTACT TCGGGCGGAT GTCCATTGGA GCCATTAGG CTAGAAAGGG CCCAGCTTA GATAGAACCC GTTGCATATG ACCTTCTC-CT TCTAACGAAA TCTAACGGTTC ACGGGCCCCC CCTCAGCGAA CATGTCCCAGC-C-ATTACTACA AGAGGCATGT TCGCAAGATC CAATTATTGG TGTAAACCTG GATT-GGTGCC TAATGTATGT GGCGATTAA CTTCGTTCT TGCTTGCTGA GTGTGGTCC-C CCCCGACTTC CGCGCGAAGT CAGATATCAA CATACTGGGCC CCTGTGCCAC CA-CTGGTGAC GTAATTAGAG GTCTAAGACT TTTGTGTTAC GCTCTCAGTT AGGGCC-GACG CGGCATTAA ACTTGGGAGG ATCGACGCAT CTATATCTGG CTCCTAATA-C CATATTAGAA TACACAG ATGGTGCA 6-11: ACAAAATGA TCACGCAAGG CAACC-ACGGA CCCAAATTGC GAACCACCTC TGGCGAAGAC AAACATACAC AATAATT-CCA TTCGCATGAA ACCCGAGCTT CGCGCCGAAT TAGCTGTAAG AGCAAAAATA

ACGCAGGCAG AGGGCGTGT A CCGGGGGCGT CCGAATTCTG GAGGCAGAAG T-A-GAGGTATG TCGAAAATCC ATTACCTCCG TATATTCACT TCCAATAAA TAAGG-AATCT AAAACTCGCA CCACGTGCTT TTTAGGTCTA CATAATGTGGG CCGCCGAAC-G ACCTGGTTCG TAGAATCGAA GTCGTCGCAG CAAAGACGCT CTGTT CGGGG A-TGGTAGCAG GCCCTTATA CGGTACTTT ACTTAGGGGG ATATTGGCCG AGATG-AGGTC CTGGCTTAGC ACCCCTAAAAA GCAAACACGT GGCGGAAGAC AGCAAAC-AGG GGGGTCCCAG TGAGATCCCG GGACTGCGTT AAGCGGGTCC CGGACTATC-A ATCTAGTAAC CCCTCGTA

7-8: GCATCGAC GTTGTGTTGCC AGCCGGAAAT CAAGGCAAAA AAGTTAGAG-C CTGTATGTCA GTGTTATTCTG TTATAT CTGC CTAAGCTTTT CTCCTTCGTA TGTC-AAGATC CGAATAACCA AATGATTAC TAAGTCTTCG CCGTTCTAAC ATGCTCCG-CT TGACGTGAAC TTAGGCGTCG AACGAGTTGG CGACAGAGTG GATGGTTACT A-TCAGTGTGT AGTCGTCGCC CCGTAGGACC CCATCATCAA TCCTCCCCCTC GCGTA-GCGAA CGCACCGTCG GGGAACGGGG ATTGCGGAAG CTTGCAGCCT AAATGCA-TCT AGGCCATTCA CGAAAGATGG GGAGATGTAA ATGGCCTCAA CACAGTACC-T GCATTGTGCT AAGGCCAATC GTCATTGGAA TACTCGTAAA GGCTAACGAC AG-TGAGCAAC AAGGATTAA TCGGACCTT GTGCTGTGAC AATCTGAACG TTTCAA-GCAA GAACGTAGGC GCGTCATGGA TTAGGTCCGA CTGTTCTATC GTACGCCGA-T GTGAGTGGAA TTACGCCAGG ATGGCAGGCG CGAGCCGAAT TACTCGTTAC GC-GATCCGTC CCAAATTGAT TGAATAACCT GCATGTAGAC CCACGTCCCTC TCGGCA-AGCC TGTTACGTGT TATATCTTTC TTTTCAGCAT CCTAGACAGA TCGTTAACCG T-ATAGGGATA CTACAAACTC TGATTCATCA AGACCGATGA AGTTGCTATC CCAC-GTTGCA CATATAAAGA GTTTCAAAGA GGCCCAACCT CTTCCCTCCGT ATTGCGC-CA GTAACGGTCG TTTACGCCG AATGTAATAT TGTCCCGCGT CCTACCACGG GT-TTACTCA AATCGAGCAC CGACTACGTC AACAAAGCTA ACTATGTGGA TAATC-ATAGC GACGCCAAA GATCATAGTA TTAACTCCTT AACTTTTGC GGTGTCTGA-A CACAATTGGG ATACGACTAT GGAGAGGCTC GGAGCGGAAA ACATAGCGG A-TGGTGCA 7-11: GCATCGAC CAATGCATAT ACCTAACCTA AGGCCGATAT CCCGT-CTGTT CCTCAGCCCT GGCTGACCTC CGCCCTGCTG GAGTTGCAAT TGGCACTAA-T CCTTCATCGG AGAGAACGGG CCCGGGTGT A GTATAGATGC GCTGCGAGTC AC-AGGCTGCC CGTCGTAGAT GACGATCTTA TATATAGTCT TTCAGTGAAC GGTCG-CATG TGAGTTAGAC CCGTTTCAGC GCGCGGCACA TCAGTTAGA ATGGCGCCT-T TGGCTCGTAA CCCAGGTTGA GAGTATACTC AGGTTTACA ATTGACATC TTA-ACTGATA CGTGATGGAT TGCCCGCGT TTTCGTTAA GCTCGATATT TCACGAC-TCC TATACTAGTT CGAAATTCAA TCACTGGAA GGGTACTGGG CATTTCCTAT T-ATACCTCTA AGAAGTACGC TAACAGGGC AAGTATTGGC CTACAGACTT TAGG-CGCCIG ATGACAAGCC GACTCCCCGT TTATTTAGGA CGCAGAAAGAG TTAAGTC-ACT TCCCCCTCAT CTTAATAGT TATGATTGAC ACTGAGCACT TTACAGCGGT T-ACCGAGTTA GTGAGATACG GGAATACTGC TATCTTGGT GTCTGGCG TTGAC-ATGCC ACCCCGACTA TGACAAAAAA CCATAGAAAC TAAGGAAGCA TGTGTTT-TGG GACGACGGCA TCGTTGTGGC TCCATATCGT TCTTCTGTA GATAAGCTGA GG-CCCAGCGG CACGGGGTAG TTGGGGAGTA ATCCCAGTTC ACAGTACATC GGTTCT-CTGG ACACGATTGC CCTGGCGACG ACAGCTACA AACTCTGGCT TCAGAACGG-G CTACTCCCTT CCGTAGTAAC TTACGTTCCC TCCGCGTGGG CGAGCCGAAT GAG-TAGAATT TTCCCCGTAG ACAAAAGCCTG TGAGCTGGCT TACAGTACA GGCTTTC-GCT ACTTGATGTA CGATTCTTT AGGAGCCTCA GTTCGGAGAG TTAACTTAT CC-CTCGTA

8-11: CCTACTCC TGAACAAACCA TCAGCGATAC TAATTGGCGT GTGTCTCG-TA TGAACACCTG GCTCATCTAC CTCTTT GTTA GAACTATAAT AAGGTTCTGA-A-ATCTGTGTC TTACGCTCCT CGTCGCCGAT TAGGCCGGAG CAGAATCGGT CGAA-TT CAGA TCATTTCGAT ATATGACGAG TCCTCCATGC TACAAAGAGC GCATGGG-GCG AAAATAACCCG ATAATACTTG GCCAATATG GGCGCTGACT AAAAGTTGA-G GTGGGACCGC CCCCTGATA TAGCCCGACG CGGATAAAAT GCCTACCTAT C-TC GCGAGCC TACATGCCG TTTGGCGAAT TTAACGCCAT TTAAACTGAA AAAA-CATAAC AGCCCCGGTG ACGGATGCTG GA ATCAAGGT AAATACGCAA TGCGAA-CAGA TAGCCCTCAG CATGACCGCA GTGCGTACCC CCCGAGTCCA ATCCTGCTA-T TTATCGCTCA GAGCACCGTC GTCGGCTAAC CACTCACTTG TCAAAATCGC AA-CACCGATA TATCGTGCCTA CCTATACTCA ATTACACACGG TCGTTCGCGG ACTCGC-TGTT GCCGGATTGC GTGTGGGATG CATCGGGTAC TCAGGCTACCG TAAGGACAG-T GTTCTCCCTT TCGGACCGCT TTTAGAGACG CTACAGCGGT TTAGTTGGAA GCA-CGGCCTC CGACTCCGGA CCCTCTCAT AGGAAGGTGT TCTTTACTT CTAGAACT-GA ATAGGCAGCT TAAAATTAG TTACAAGCAT TTCGGAAATG GCTACGTAAC A-GCACAGGAC TTATACGGCG CGGTAGTCGG AAGTAACGCT TTGGTCAAGG CCCC-ACAAAA GCAACCAAAC CAGAAAAC AG GACGAGATAA GTTTTCCAT CGAATG-TAAC GGAATGACCT TAGTTCGCTT AGGTGATGAT CGCACGAGTC TACGATAGT-A TGGTGGGTAC CACGGCCGGC TACTCAGGCC CCGCACGGCG GAGTGGATGT C-ATCAATGA CCCTCGTA

9-10: GGTGAAA AAAAGTGTGCG GCATCGTCCT AGCAAACAGG CCTACTCG-TG TTTTACGTCC AAAGGAAACG CACT GGACAT AGAAAGTGCT CCTTTACGA G-ACATGCGTG TCTTCCCACG ATAGTCCCGC GGGAAAGCTCA CGGAACCTAG GGTC-TGTAAC AAATTCCGTA ACTCGGCTAC CCGAGGACAC GATTACGAG GGGCGGT-GGC CGGGGAAATA GCCCCACCAC GGCCGGGGAC CGGGTTAGTA CTCGTGGCG-A GGGAAAGGGCT CCAAAAGCCG TCGATCGCAC GATTCCGTCG GTCCGAGTCG C-AGAAGGAGA CCAGACAGTC AGCGGCCGTA CCCAGAATGG CCAAAGCAA GCC-TAATTGC AGCACTGAAA GTAGCGGCCG GCCACCTGTT GGGGGATCTT ATTCCT-GTT GTCTGTCCCTC GGAAATCAGA TTATGGAAAT CAGCATTATT CAAACACTTG C-CTCAACTGGA ACACGGTGT A TCTCTCATCA TGACCGAAAA CGGGCTTTA TACGT-ACATG CATATCGCGT ACCCACATT GACGTAGTAC TCGTCTCAA TTCGTGGTT-C GACTTGTAAAG CCAGGCAGAG CCGCACCCCTG GGTGTCAAGT TGCTGAAACT GG-CCACTGCT AGCATATGCA CGAGCTTGT A TCTCTCTCA ACCACCGGT CCGCGG-GGAG CAACGGGGA T CCTCGCTAAC TCAAGGCAAT TGCGGGGTGA GGGCCAGG-CG TGACCTGGGT AGAGAACACA TTCACCAACA GAGCGCACAT CACATCATTA T-ACTTGTGG TACTTCTAA AAAATTGCAC GGCCTCCCT ACGTTAGTAT CTAGTC-TTGA AATCTGTGGG CAGGCCGGGT GATCAACCCG TTCTAACCGA GCGAGATGC-T AAAAGCATGA TTCACATTCT AGGGCAGGAA GGTATTGTT ATAGAACTCC CT-GGTGCGTCA AGAAAAGATG TAACACTATC TGATCAAGCG ATGGATGAGC GATGG-TATC ATT TGCCA.

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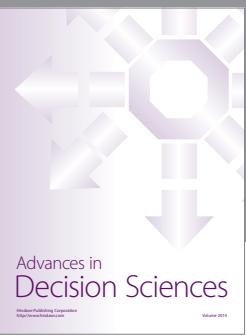
Problem. We are also indebted to Dr. Qi Feng and Dr. Xue Jie for help in the design of DNA coding.

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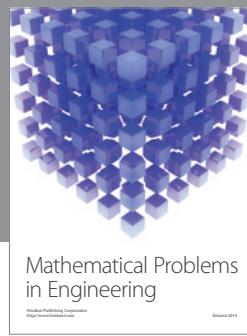
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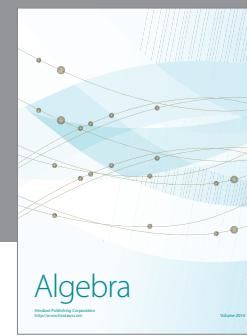
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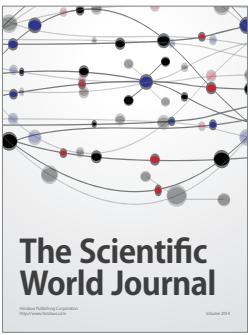
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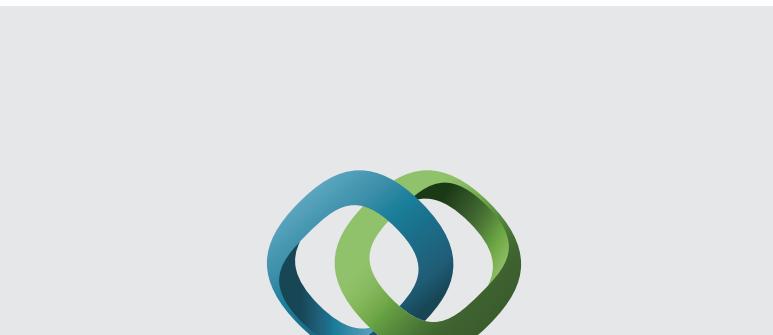
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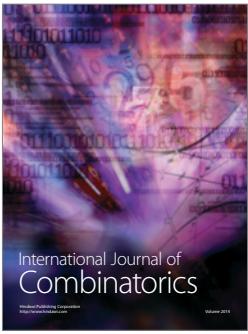


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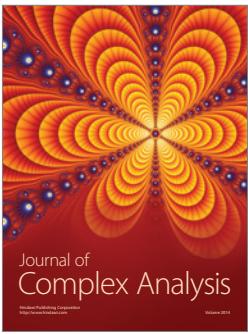
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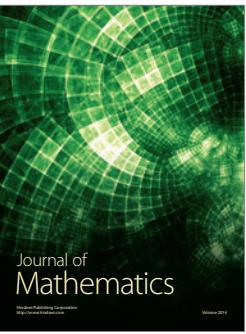
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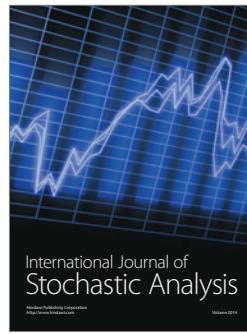
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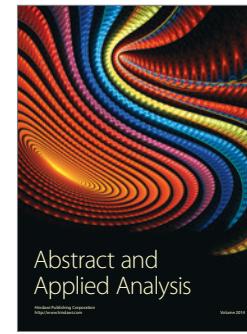
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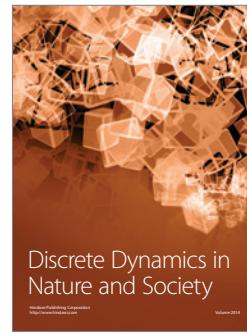
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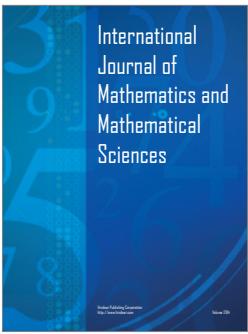
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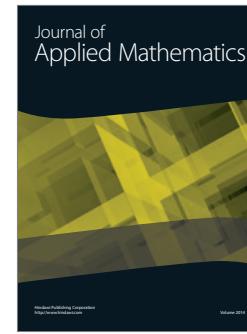
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