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Abstract: Chromosomes are essential genomic information carriers. The identification of chromosome abnormalities is an essential part of diagnosis and treatment of genetic disorders such as chromosomal syndromes and many types of cancer. There are different types of chromosome aberration like mapping of the common duplications, deletions, and translocations. One of the most considerations in the field of chromosome analysis is the segmentation. Currently available cytogenetic imaging software is designed to classify only normal chromosomes [9]. The automation of chromosome analysis is involving segmentation of chromosomes and classification into 24 groups [8]. The segmentation and overlap of chromosomes are a major step toward the realization of homolog classification. Resolving chromosome overlaps is an unsolved problem in automated chromosome analysis and the analysis is complicated by the occurrence of clusters of overlapped chromosomes. Current systems for automatic chromosome classification are mostly interactive and require human intervention for correct separation between touching and overlapping chromosomes, and so are unable to segregate overlapping chromosomes. In this paper, propose an automatic procedure to obtain the separated chromosomes. The overlapped chromosomes are segmented by found the interesting (concave and convex) points with the help of chain code algorithm and the interesting points [1] are located on contour of the image and then the curvature function is found to find out concave points and the possible separation lines are plotted by using all concave points and finally construct hypotheses for possible separation lines between concave points. The segmentation is carried out by means of a curvature function scheme, which proved to be successful.

Index Terms: Diagnosis, genetic disorders, automatic chromosome classification, interesting points, hypothesis.

I. INTRODUCTION

Fig 1: Touching and overlap chromosomes

CHROMOSOMES are microscopic structures containing all of an individual's genetic information. The chromosomes are composed of DNA and various proteins. The DNA contains the actual genetic code of an individual, and the proteins protect the DNA and allow the DNA to duplicate properly when the cell divides. Chromosome analysis is performed on dividing cells in their metaphase stage [9]. The metaphase stage images are shown in fig. 1. The chromosomes in a metaphase image may be bent, their arms may be joined (along points other than the centromere), their bands may be spread, and they may touch or partially overlap each other.
Thus, all these problems should be taken into consideration when processing the metaphase image. Current systems for automatic chromosome classification are mostly interactive and require human intervention for correct separation between touching and overlapping chromosomes. The separation is performed by posing a series of hypotheses that fit certain constraints [13], [17].

II. RELATED WORKS

1) Segmentation methods try to separate between touching chromosomes by classifying the chromosome pixels and the background pixels into two different segments. Since segmentation methods are general methods that do not depend on the shape of objects, their link to the specific problem of chromosome separation is weak, and, therefore, they tend to fail in cases of incomplete information. A case of incomplete information occurs; there is no separating path between chromosomes [5], [7].

2) Heuristic search edge-linking methods try to separate between touching chromosomes by searching for a minimal-cost connected path that separates between the chromosomes. The link lost connection of such methods to the specific problem of chromosome separation leads to inferior results in cases of incomplete information, such as in the case where a separating path does not exist [19].

3) The method for separating the touching chromosomes by using shape decomposition based on fuzzy subset theory [16] is reported as giving inferior results relative to other general methods. An even greater disadvantage of this method is that it also yields erroneous decompositions of single chromosomes in cases of bent chromosomes.

III. PRELIMINARY PROCESSING

The preliminary processing of the metaphase image is performed by increasing the effective sampling resolution [14] may be achieved by sampling each metaphase image in some overlapping parts. Fig. 2 is the sample overlapped and touching image.

A. Obtain the Chromosome Contours

The relevant information of a shape is contained in its contour, for separation purposes, only the shape contours are required. The use of shape contours actually represents a successful information compaction that reduces the amount of processed data and stresses the main features of the objects. Chromosome contour determined by using a common contour-following method to track the contours of the connected segments by row and columns [2]. Fig. 3 shows the binary approximation of the overlapped image and fig.4 shows the contour of the overlapped image.

Fig 2: Sample overlapping and Sample touching images

Fig 3: Binary approximation of the given overlapped image

Fig 4: Contour plot of the above binary image. Contour plotted in green
B. Obtain the Discrete Curvature Function

The curvature function of a curve is defined as the rate of change of the curve slope with respect to its length. The radius of a circle is tangent to the concave side of the curve, and has the same curvature as the curve at the tangential point. The curvature function is a derivative of the contour’s slope function. To reduce the computational requirements of the K-slope evaluation, an extended chain-code method is used. Using the extended chain-code, it is possible to determine the K-slope at a given pixel directly from a look-up table where the extended chain-code serves as the entry point.

This method used for detecting and locating "corners" in chain-coded curves [3]. The "corner" means that chain node with which we can associate an identifiable discontinuity in the mean curvature of the curve. The detection of a corner is a function of the magnitude of the discontinuity. The concave and convex points of the object are detected by using chain code method. Fig. 5 presents the result of interesting points detection on the curvature graph, where the interesting points are marked with triangles.

C. Obtaining the Interesting Points

The interesting points are detected as extremum points of the curvature function, or as middle points of constant curvature curve segments. The required filter should combine a low-pass filter for high frequency noise removal, and a high-pass filter for curvature peaks enhancement.

Fig 5: Interesting points on the curvature graph.

Fig 6: Interesting points on the chromosome

Fig 7: Extended view of the above chromosome image

Thus actually resulting in a band-pass filter. The filtered curvature function is segmented, and extremum points are detected within each segment. From that list, only concave points with a curvature measure above a certain threshold are selected. The interesting points (concave and convex points) are located on the chromosomes. Fig. 6 shows the location of interesting points on the chromosome and fig. 7 shows the extended view of fig. 6.

IV. CHROMOSOME SEPARATION

Each chromosome as a separate subset of pixels, it is possible to view the chromosome separation, the minset is obtained by taking the intersection between some subsets and the complement of the other subsets. Each minset of the chromosomes represents either a part unique to one chromosome or a part common to more than one chromosome. The minset normal form representation of the chromosomes can be constructed by a union of such parts. The required partition is obtained by splitting the object along lines that connect high concave points.

For the determination of separation lines [4], all the possible pairs of high concave points are considered. The separation lines are obtained by taking all the possible combinations of two points.

A. Hypotheses for Overlapping Objects

The constraint requires that the spatial distance between the vertices of a separation line of touching objects [20] should be small relative to the distance between the vertices along the shape’s contour.
To construct a hypothesis for the separation of overlapping objects, an overlap minset should be located. The distance between the separation lines should be above a given threshold. Fig. 8 shows the possible separation lines between minsets.

**B. Hypotheses verification**

The verification of hypotheses is based on an evaluation of the fit of the obtained parts to the prototype shapes of the chromosomes [10]. In general, the shape of a chromosome is not fixed. A chromosome may have various heights and widths.

The evaluation of the fit of a chromosome to its prototype shape is done by fitting a bounding polygon to the chromosome [10], [11]. The bounding polygon is constructed by locating the shape’s minimal bounding box, contracting it in the middle, and rounding its corners. In order to retain the uniqueness of the bounding polygon as a chromosome prototype, constraints are posed on possible locations of its vertices. The minimal bounding box of a shape is found by computing the principal component transformation of that shape. Fig. 9 shows the overlap region, fig. 10 shows the first chromosome segmentation from other chromosome and fig. 11 shows the second chromosome segmentation from other chromosome.

**V. FUTURE WORK**

In the case where there are only touching objects, the problem of finding a separation hypothesis is reduced to finding the minset partition. After finding that partition, each object is composed of one minset and therefore, there is no need to find unions of minsets. Since the number of possible partitions is rather large, it is reduced by introducing an additional constraint on the possible pairs.

The constraint requires that the spatial distance between the vertices of a separation line of touching objects should be small relative to the distance between the vertices along the shape’s contour.

The reduction in the number of possible partitions by using the constraint of nearly parallel separation lines. An additional reduction of the number of possible partitions is obtained by introducing a constraint on possible combinations of pairs and so possible partitions will be only part of the sets.

For example Fig. 12 shows touching region and fig. 13 and fig. 14 shows the disentangling of touching chromosomes. The experiments are going in the separation of touching chromosome with out any loss in the information [12].
A. Sample disentangling Images

Fig 12: The region of touching chromosomes

Fig 13: Disentangling of first chromosome from second chromosome

Fig 14: Disentangling of second chromosome from first chromosome

V. CONCLUSION

Karyotype analysis is a widespread procedure in cytogenetics to assess the possible presence of genetics defects. The procedure is lengthy and repetitive, so that an automatic analysis would greatly help the cytogeneticist routine work [6]. Still, automatic segmentation and full disentangling of chromosomes are open issues.

The problem of extracting individual objects from overlapping configurations is one that occurs in a number of applications in machine vision [18]. It is a difficult problem in general and can be thought of in two parts: 1) identifying overlapping objects in the first place and 2) identifying the individual components. In the case of chromosomes, it is usually fairly easy to identify an overlap on straightforward shape criteria.

This proposed method for Automatic Chromosome segmentation of touching and overlapping images are more accurate than other previous methods and this method of segmentation is applicable for large number of bents present in the chromosome structures and more number of clusters [15] by the location of each separation point is optimized, yielding the minimal possible distance between the smoothed approximation and the original curve.

The experiment proved that this algorithm can segment the overlapping chromosomes successfully.

REFERENCES


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