



Congenital Malformations Ultrasound-Based Fetal Prediction using a Computer-Aided Diagnosis System

W. Fathima Farsana^{1*}, Dr. N. Kowsalya²

^{1*} Research Scholar, Research Department of Computer Science and Application, Vivekandha college of Arts and Sciences for Women, Elayampalayam, Tiruchengode, Periyar University, Salem, India.

² Assistant Professor, Department of Computer Science, Sri Vijay Vidyalaya College of Arts and Science, Nalampalli, Dharmapuri, India.

*Corresponding Author Email: afsheensyed84@gmail.com

Abstract

Notwithstanding all of our current knowledge, cutting-edge diagnostic tools, computerised databases, and other readily available supporting resources, fetal syndrome detection continues to be a challenge for healthcare professionals both in prenatal and postnatal periods. The early treatment of fetal disorders is a difficult puzzle to put together and solve. One anomaly should always raise concerns about the existence of everyone else, and it may act as a spur for further research and increased awareness of other disorders. The measurement of the standardized plane is necessary for biometric assessment and diagnosis during an ultrasound (US) examination. Based on a review of existing algorithms for autonomously monitoring fetal development, a unique genetic algorithm named Neuro-Fuzzy was developed. First, the benchmark image from the fetal ultrasound is automatically pre-processed to use the Normal Shrink Elliptic curve method. Second, to extract features, Rotation Invariant Moments (RIM), Intensity Histogram (IH), and Gray Level Co-occurrence Matrix (GLCM) are utilized (IM). Finally, a genetically based Neuro-Fuzzy approach is used to distinguish between abnormal and normal prenatal growth. When compared to state-of-the-art approaches, experimental results using a benchmark and actual dataset reveal that the suggested strategy achieves 97% accuracy in terms of parameters like Sensitivity, Specificity, Recall, F-Measure, and Precision Rate. The Support Vector Machine (SVM) achieved the highest accuracy rate of 95.7 percent in comparison to certain other classification methods such as KNN, Ensemble methods, Linear Discriminate Analysis (LDA), and Decision Tree, the with ROC curve covering 0.97 SVM. The area underneath the receiver of characteristics (AUC) and the confusion matrix are also used as assessment indicators.

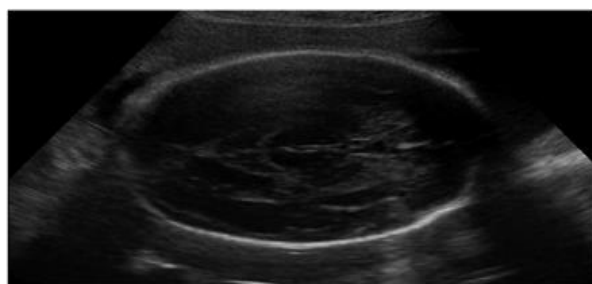
Keywords: fetal syndrome detection, ultrasound, Rotation Invariant Moments, Intensity Histogram, Gray Level Co-occurrence Matrix, Neuro-Fuzzy approach, Support Vector Machine.

1. Introduction

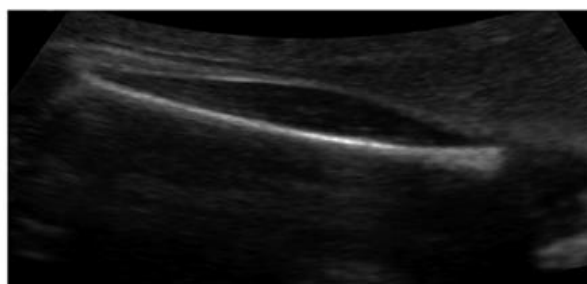
Ultrasound imaging is frequently utilized since it is less expensive and non-persistent than other imaging techniques. Image properties such as Echogenicity, Legion Shape, and Echo Texture are used to diagnose a variety of disorders [3]. For the delivery of high-quality

obstetrical health care, accurate ultrasound-based fetal biometric assessments are critical. The Bi-Parietal Diameter (BDP), Head Circumference (HC), Abdominal Circumference (AC), Femur Length (FL), Humerus Length (HL), and Crown Rump Length are all common measures (CRL). Guidelines for assessing these values are published by the American Institute of Ultrasound in Medicine (AIUM). Growth restriction, microcephaly, and macrosomia are among the fetal pathologies that these readings may assist identify. Furthermore, they are used to calculate the fetus' gestational age (GA) (i.e., length of pregnancy in weeks and days). A ultrasound will reveal a fetal structural abnormalities in around 3% of pregnancies, which may vary from a single minor flaw to severe multisystem malformations that are lethal[4].

In the examination and clinical triage of prenatal structural malformations, genetic studies are critical. Traditional prenatal cytogenetic analysis was the gold standard for investigating congenital aberrations for more than 30 years, but in the past decade, chromosomal microarray analysis has been more widely used to discover submicroscopic harmful copy number variants (CNVs) in prenatal diagnosis. (5), (6) When chromosomal microarray testing is added to karyotyping, the prevalence of chromosomal abnormalities is increased by 3–5%. [7] Aneuploidy, uniparental disomy, CNVs, and intragenic mutations are all examples of genetic variation that may cause fetal anatomical abnormalities. Genome-wide sequencing techniques to analyze prenatally diagnosed congenital disorders are gaining popularity. WGS (whole-genome sequencing) during pregnancy has previously been documented [8]. Whole-exome sequencing (WES) and targeted gene panels, on the other hand, have attracted increased attention because to their cheaper cost, the smaller quantity of fetal DNA needed, the prospect of a faster turnaround time, and better sequencing depth. [9, 10, 11, 12] We previously employed WES in 29 fetal-parental trios with a fetal structural abnormality, and in 10% of those instances, we found an underlying genetic etiology. 7 WES has shown variations in genes linked to developmental abnormalities in more than half of the fetuses in studied cohorts, although most prior investigations [12, 13] have employed small sample sizes (of 30 fetuses or less), are limited to highly chosen subgroups (eg, those obtained at autopsy), or both (appendix).



(a) Fetal head (28 weeks)



(b) Fetal femur (28 weeks)



(c) Fetal abdomen (28 weeks)



(d) Whole fetus (13 weeks)

Figure 1: Baby ultrasound images of the head, the leg, and the abdomen reveal a fetus at 13 weeks of gestation.

The Prenatal Assessment of Genomes and Exomes (PAGE) research was developed to investigate the use of genome-wide sequencing techniques in the prenatal identification of fetal structural abnormalities. We aimed to report the WES results from 610 fetuses (and parental samples) with a wide range of fetal structural anomalies as part of this larger project, and to highlight the ethical and practical issues we encountered that have implications for the translation of prenatal WES into clinical practice.

Routine ultrasonography detects fetal congenital abnormalities in 2 percent to 5% of pregnancies [14, 15]. The presence of congenital defects may be distressing for expecting parents and have a considerable influence on perinatal mortality and long-term morbidity [16, 17]. The underlying etiology of these abnormalities is complex, and genetic factors play a role. Molecular rapid aneuploidy testing (RAD) and chromosomal microarray analysis (CMA), which are used to identify numerical and structural chromosome abnormalities, respectively, have a combined diagnostic yield of around 40% [18, 19]. However, this implies that the underlying cause of the observed congenital defects is unknown in the vast majority of instances. Congenital abnormalities induced by monogenic diseases caused by point mutations and/or minor insertion deletion events are particularly prevalent in this category. Depending on the clinical preselection of the cohort and subsets of genes investigated, whole exome sequencing (WES) in a postnatal scenario has been demonstrated to boost diagnostic yield for genetically heterogeneous (monogenic) illnesses by up to 58 percent [20, 21].

Regular WES turn-around times (TATs) of many months have always made it difficult for this assay to be used in routine prenatal diagnosis. Reduced TAT may aid in the identification of babies with congenital abnormalities for whom a genetic diagnosis was elusive utilizing standard prenatal methods. Rapid whole exome sequencing (rWES) has been proven to aid clinical decision-making in pediatric and neonatal critical care [22-25], with TATs ranging from 4 days to several weeks. It's extremely conceivable that rWES can help with prenatal clinical decision-making in the same way. 81 percent of cases were genetically identified in recent research on the use of rWES for fetuses with skeletal abnormalities. 14 Although this rise in diagnosis allowed for more accurate pregnancy outcome prediction, giving parents greater confidence in prenatal decision-making, skeletal defects account for only around 30% of all fetal congenital malformations [26,27]. In a few pilot studies [28-31], the effectiveness of using rWES as a first-tier test for the whole spectrum of prenatal congenital abnormalities found during regular ultrasound imaging was also investigated. The great majority of these investigations focused on diagnostic yield and TAT as outcome metrics, rather than the impact of the rWES result on clinical decision-making. The rWES findings of 54 fetuses with congenital abnormalities in continuing pregnancies are presented here, with an emphasis on the impact of rWES on clinical decision-making.

According to the European Registry of Congenital Malformations (EUROCAT), the prenatal detection rate for 18 congenital anomalies excluding genetic conditions ranges from 44.8 percent for clubfoot (talipes equinovarus) to 98.4 percent for anencephaly and similar conditions, while all genetic conditions were detected in 72.8 percent of cases, ranging from 66.1 percent for Down syndrome to 93.6 percent for Edwards syndrome. In the EUROCAT [28], the total detection rate of all anomalies was found to be 34.5 percent. There are no data on the prenatal detection rates of several disorders. If at least two congenital abnormalities have been found prenatally, a fetal syndrome should always be looked for and evaluated. All

kinds of acrocephalopolysyndactyly have a low prevalence rate, ranging from 0.02 per 10,000 births to 0.96 per 10,000 births for DiGeorge syndrome [29].

The purpose of this study is to show and debate the potential of several US methods for improving prenatal detection rates of particular fetal disorders.

2. Literature Review

The study of ultrasound imaging has progressed significantly, and the application of contemporary machine learning in the medical image analysis sector has made this art more difficult for researchers [1]. Computer Aided Diagnosis (CAD) has become a hot topic in medical imaging and diagnostic radiology research. The rationale and philosophy for early creation of CAD, as well as the current state and future possibilities of CAD, are given in this research article. The overall research framework advocated that more research and development be done. This framework was inspired by traffic sign detection and is now being developed and used to the diagnosis and development of liver cancer. Ultrasound imaging is a prominent and commonly utilized modality for seeing and analysing medical pictures for any illness state without giving the patient any pain or suffering [2].

Prenatal identification of congenital abnormalities became possible with the development of ultrasonography (US). Despite this, detection rates in ordinary clinical practice remain inadequate. Some writers suggest that three-dimensional ultrasound (3D US) improves the identification rate of all abnormalities, whereas others remain unconvinced. Only a few hundred and counting syndromes may be diagnosed prenatally, according to current knowledge of the prenatal detection rate of more than 6000 syndromes.

Prenatal detection, on the other hand, is far more accurate and precise when new concepts and information from scientific study are combined with advances in diagnostic technology. There has been a huge advance in US equipment, with outstanding picture quality, thanks to the current dynamic and rapid growth of computer technology, such as 3D high definition live (3D HDlive) silhouette and flow US technology. The use of ultrasound to test the functioning of various organs and systems, such as the brain and eye, has given rise to new disciplines of fetal evaluation, such as fetal neurology and fetalsonology-ophthalmology [30]. Some innovative functional tests, such as the Kurjak antenatal neurodevelopmental test (KANET), have been incorporated into routine clinical practice to examine the function of the fetal brain [31–34], providing further useful input into the diagnosis of fetal disorders [35]. Prenatal identification of fetal syndromes has shifted from the second to the first trimester of pregnancy, according to several publications [36, 37].

The fetal ultrasound picture segmentation was described in the paper as a semi-automated approach for estimating fetal weight (EFW). The described technique measures four homogenous fetal parameters: abdomen circumference, abdominal circumference, head circumference, and biparietal diameter. Support vector machine, regression analysis, and neural networks are some of the classification techniques used to assess EFW, head, and abdominal data. The main objective is to develop a period-assured morphology-based computation to sense femur form in fetal ultrasound images, modify its shape for programmed length measurement, and achieve accuracy and consistency of estimation in this manner [38]. The images obtained from the individuals were initially processed using morphological administrators to remove the background from the image. The images were then changed using morphological administrators to improve the condition of the femur until a single pixel – wide skeleton of the femur was available in the most time-effective manner. The femur end-focuses are supposed to be the skeleton-end-indicates, and the femur length is

calculated as the distance between the end-focuses to estimate gestational age. The suggested computation has been tested on real clinical photographs, and the results have shown that the estimates produced by the proposed method are accurate and in good agreement with the manual estimating technique. [39] investigates two ways for division groups of fetal ultrasound images: one is location development, and the other is a version of split and consolidation computations. They provide a simple architecture for processing and segmenting a large number of highlights fast. The UI was created using the Tcl/Tk toolkit, which is freely available. The framework's most serious flaw is the lack of credible metrics for evaluating the division's precision. It also necessitates systems that are more tolerant of disturbance and antiquity, since district development planning is quite sensitive to the surrounding community. Another option is to depict the limit using a Binary Space Partitioning tree (BSP). [40] proposes a fully programmed division that distinguishes ordinary and ultrasonic tumor liver images using factual highlights.

The speckle noise in ultrasound images was separated using a pinnacle and valley method, then the image was smoothed and a moment organize channel was used to channel the turmoil and improve the picture's quality. The peak and valley channel suggests an interesting replacement for the central channel with the purpose of increasing efficacy. [41] describes the use of contourlet change for de-speckling ovarian ultrasound images, dynamic forms without an edge technique for division, and fluffy reasoning for characterisation. The experiment was carried out using test ultrasound images of the ovaries, and the results were compared to the deductions made by an interim based classifier as well as those made by experts. [42] shows the extracted image highlighting the use of different computations that are indicated with developing models with inner modules. This project actualized an unique FPGA-based engineering enabling continual extraction of four GLCM highlights. For inclusion extraction, a 128 By 128 dim level image was obtained.

Using the measured second request approach, the digital image highlights were extracted. We were able to extract an image in dark scale level evaluation using this measured approach. Several essential highlights of ultrasound kidney images were separated in this study, including the Intensity Histogram (IH), Invariant minutes (IM), Gray level co-event grids (GLCM), Gray level run length networks (GLRLM), and a 'Consoli-dated' list of capabilities created from a mix of all four highlights [43]. A total of 48 highlights from each image were identified. In the preparation of datasets, the consolidated element acquired a perfect accuracy rate of 100 percent. It was successful in organizing 94 instances from 94 occurrences. It correctly sorted 35 out of 40 instances in testing datasets, with an accuracy rating of 87.5 percent. This revealed that the combined element's usage was somewhat effective. [44] has tested with a strategy suggestion for arranging liver ultrasound images in light of surface examination. It makes use of a set of seven surface features with strong discriminative strength that radiologists may use to describe the liver.

[45] discusses a hybrid strategy for automated identification and measurement of belly shape from 2-D prenatal ultrasound pictures that combines classic texture analysis approaches with deep learning. When the CNN projections were combined with those from the slope boosting machine (GBM) using histogram of situated inclination (HOG) highlights, a notable improved division was obtained.

3. Methods

3.1 Syndrome Prenatally Detection

The nomenclature used to define fetal disorders might be misleading at times. There are many terminologies and synonyms used. There aren't always clear definitions of how many

main and minor criteria are required to identify each condition. Discrepancies in screening strategies and follow-up methods, as well as probable disparities in practitioners' abilities and accessible equipment, may all contribute to differences in prenatal detection rates by area or nation.

Clinical dysmorphology is a discipline of clinical genetics devoted to the study of aberrant human development, with a focus on disorders shown primarily as morphological changes in the body. Fetal maldevelopment may be classified as malformation, distortion, disruption, or dysplasia due to a variety of pathophysiological processes. Malformation is typically characterized as a single localized fault in tissue formation that sets off a chain of defects (e.g., anencephaly). Malformations have a recurrence risk that varies from 1% to 5%. Extrinsic mechanical stresses strain ordinarily normal tissue, causing it to deform (e.g., abnormal faces, pulmonary hypoplasia and limb contractures that result from prolonged oligohydramnios or primary renal agenesis in Potter syndrome with Potter facies). An external injury causes disruption by destroying normal tissue and changing the development of the afflicted structure (e.g., amniotic band syndrome). We talk of dysplasia when the major problem is the lack of normal cell organization into tissue (e.g., achondroplasia). If many organs are implicated, any of the causes of prenatal maldevelopment may result in abnormal morphology of fetal organs and systems, leading to the establishment of a fetal syndrome. The term "syndrome" comes from ancient Greek and means "running together." It refers to a grouping of indications, symptoms, dysmorphic characteristics, and/or behaviors that recur in the same person.

Some fetal disorders may be discovered before birth, while others cannot; some fetal syndromes are manifested before birth, while others are not. Many times, even many years later, a clear diagnosis may be established postnatally. The differential diagnosis of fetal syndromes is extensive, and there are various online databases that may aid in the identifying of anomaly patterns as a syndrome, sequence, or relationship. Online Mendelian Inheritance in Man (OMIM), Orphanet, London Dysmorphology Database, Possumweb, and the Phenotip online database are the most extensively used online databases. The Phenotip online database is the most user-friendly database for sonographers, as it is specifically designed to include all antenatal sonographic findings (rather than postnatal findings), whereas the London Dysmorphology Database and Possumweb are non-free databases that aid in differential diagnosis and include postnatal findings. There is rapid access to information, with the option to search by ultrasonographic marker, a combination of a few markers, or just by the syndrome's name. Known family history, previous pregnancy with a malformed fetus/infant, history of consanguinity, exposure to a teratogenic drug or other agent, travel to high-risk areas, and possible exposure to infections (Zika virus, TORCH infections) or trauma could all be triggers to look into the syndrome further. If parental markers are present, they may also be included. Synonyms for the syndromes are automatically included in the search, making it simpler and quicker.

3.2 Pre-processing

Speckle noise reduces the image's contrast and results in ringing effects around the edges. As a result, the capacity of humans to discriminate between coarse and fine elements of a picture is harmed. There are a variety of filters that may be used to reduce speckle noise. Each filter has its own set of statistics. For each picture, a different filter produces various results. The researchers are examining many state-of-the-art denoising filtering approaches, including Wiener, Median, Lee, and Kuan, in order to enhance the performance of these filters in terms of quality, run time reduction, and other parameters.

3.2.1 Median Filter

The median filter is a non-linear approach that works well with impulsive noise (salt and pepper noise) and keeps the image's crisp edges. This filter substitutes the center pixel inside the define window pane with the middle value in the collection and types the enclosing pixels value in the window pane with an ordered collection.

3.2.2 Kaun Filter

The Kuan filter is said to be better than the Lee filter. Within the filter window, it does not make any approximations to the noise variance. It merely converts multiplicative noise, such as speckle noise, into an additive linear form, but it uses an image's Equivalent Noise Look (ENL) to produce a different weighted 'W' for filtering.

Filter Algorithms:

```
1. allocate outputPixelValue[image width][image height]
2. allocate window>window width × window height]
3. edgex := (window width / 2) rounded down
4. edgey := (window height / 2) rounded down
   for x from edgex to image width - edgex do
   for y from edgey to image height - edgey do
     i = 0
     for fx from 0 to window width do
       for fy from 0 to window height do
         window[i] := inputPixelValue[x + fx - edgex][y + fy - edgey]
         i := i + 1
       sort entries in window[]
       outputPixelValue[x][y] := window>window width * window height / 2]
```

3.3 Feature extraction and selection

Feature extraction is a technique for identifying effective strategies for presenting anomalies in medical images. After pre-processing the pictures, which is known as the data-cleaning step, classification characteristics are extracted from the cleaned images utilizing procedures.

3.3.1 Gray level Co-occurrence Matrix (GLCM)

"Spatial Dependency" is another name for GLCM. It's one of the most widely used mathematical techniques for extracting texture information from photos. The pixel intensity level of the neighboring pixel is always the focus of this approach. The pixel's relative location to another pixel is always taken into consideration by GLCM. This is a basic metric that shows how often certain combinations of pixel brightness levels appear in medical photographs.

3.3.2 Gray level run length Matrix (GLRLM)

It's a matrix that may be used to extract characteristics relevant to texture analysis. GLRLM is a 2D matrix in which component "p(k,l)" gives total number of successive operations of length "l" at grey level "k" for each given 2D picture. The letter "M" stands for "maximum

run length" (run length is considered to be a number of neighboring pixels which possess same grey level intensity in a specific direction). GLRLM Feature Selection's representation of the feature retrieved. The generated data includes numerous redundant or unnecessary characteristics since all features were taken from the picture. The features selection approach is used to get rid of the redundant and unnecessary characteristics and locate the important ones that will help with the rest of the analysis. During the feature extraction procedure, a total of 16 features were recovered from each picture, however due to the large number of features, the neural network could not utilize all of them. Although each attribute is useful in categorization, only a handful are particularly helpful in categorizing and identifying illness states. As a result, rather than utilizing all of these traits as input, only those of high relevance were chosen.

Element difference moment of order k : $\sum_i \sum_j (i - j)^k C_{ij}$

This descriptor has relatively low values when the high values of C are near the main diagonal. For this position operator, high values near the main diagonal would indicate that bands of constant intensity running "1 pixel to the right and 1 down" are likely. When $k = 2$, it is called the contrast:

Contrast = k : $\sum_i \sum_j (i - j)^2 C_{ij}$

Entropy = k : $\sum_i \sum_j C_{ij} \log C_{ij}$

3.3.3 Feature selection

The generated data includes numerous redundant or unnecessary characteristics since all features were taken from the picture. The features selection approach is used to get rid of the redundant and unnecessary characteristics and locate the important ones that will help with the rest of the analysis. WEKA software, version 3.6.9, was used for feature selection. Only '.arff' data files are compatible with WEKA and are recognized. As a result, a '.arff' file was created, which holds the value of the extracted features (including both normal as well as abnormal). During the feature extraction procedure, a total of 16 features were recovered from each picture, however due to the large number of features, the neural network could not utilize all of them. Although each attribute is useful in categorization, only a handful are particularly helpful in categorizing and identifying illness states. As a result, rather than utilizing all of these traits as input, only those of high relevance were chosen.

3.4. Classifier

In a CAD system, classification is crucial among several domains, and its effectiveness is dependent on the efficient performance of various attributes as well as the classifier selection. Computer vision, pattern recognition, and other machine learning methods are frequently used in the area of medical imaging. Here, ANN is used to diagnose fetal development and achieves a high level of classification accuracy. However, we believe that ANN has yet to demonstrate its value; as a consequence, we have enhanced artificial neural networks using fuzzy logic and compared the classification results with ANN.

3.4.1. ANN

ANN is made up of basic parts that are inspired by biological neural systems. An ANN's structure is defined by a 'input' layer, many 'hidden' layers, and one 'output' layer, all of which

are linked via a directed network with nodes known as neurons. The most practical design of an ANN is the multi-layer perceptron (MLP) network, in which each neuron is linked to a number of its neighbors, with changing weights reflecting the relative influence of the various neuron advices to other neurons. The measured weighted summation of the inputs may be sent on to hidden neurons, where it can be altered using an activation function. Following that, using the hidden neurons as inputs to the output neuron, accurate prediction algorithms are obtained.

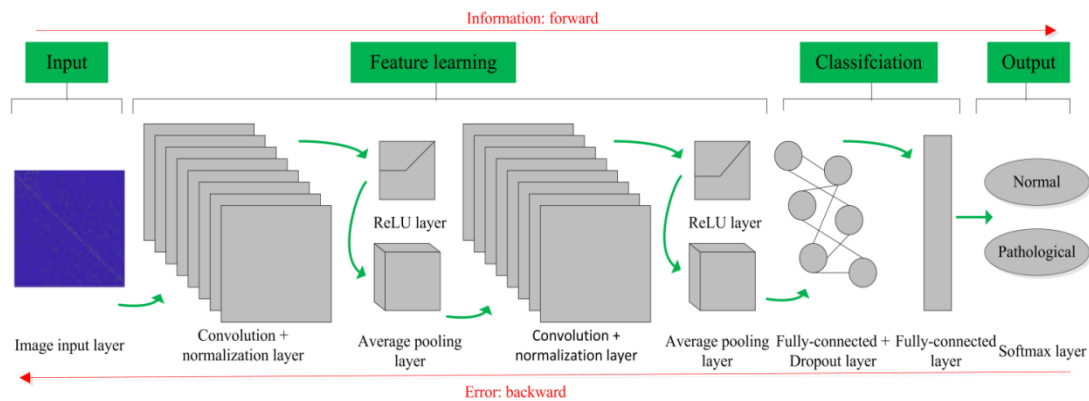


Figure 2: Neural Network Layers

3.4.2. Decision tree

In machine learning, data mining, and statistics, a decision tree is a predictive modeling strategy. It's a supervised learning method, which means it has an output variable that must be predicted from a collection of input variables. We create a function with these variables and map the input to the desired output. This method divides the population into two or more homogenous sets, utilizing major features as independent variables to create multiple separate groups wherever feasible. The target variables are represented by the leaf nodes in a decision tree, whereas the input variable is represented by a route from the parent node to the children node.

3.4.3 Linear discriminant analysis (LDA)

LDA is a technique for finding a linear combination of characteristics that describes or separates two or more classes of object or events that is used in statistics, pattern recognition, and machine learning. It's a pre-processing dimensional reduction approach for pattern categorization and machine learning applications.

3.4.4 SVM (Support Vector Machine)

One of the classifying approaches is SVM. This approach is used to "plot each data item as a point in n-dimensional space," with "n" denoting the "number of characteristics." The Maximal-Margin Classifier explains how SVM works. In your data, the numeric input variable (x) creates an n-dimensional space. If two input variables are used, the result is a two-dimensional space. The input variable space is divided in the graph by a hyperplane line. In SVM, the hyperplane chosen divides the points in the input variable space according to their class.

3.4.5 KNN

KNN is one of the most straightforward algorithms to grasp, yet it performs well in reality. It's a slow learning algorithm that isn't parametric. It's non parametric, which means it doesn't make any assumptions about the data distribution. And lazy implies it doesn't make any generalizations based on the training data points.

$$Precision = \frac{True\ Positive}{True\ Positive + False\ Positive}$$

$$Recall = \frac{True\ Positive}{True\ Positive + False\ Negative}$$

$$F1\ Score = \frac{Precision * Recall}{Precision + Recall}$$

Results and Discussions:

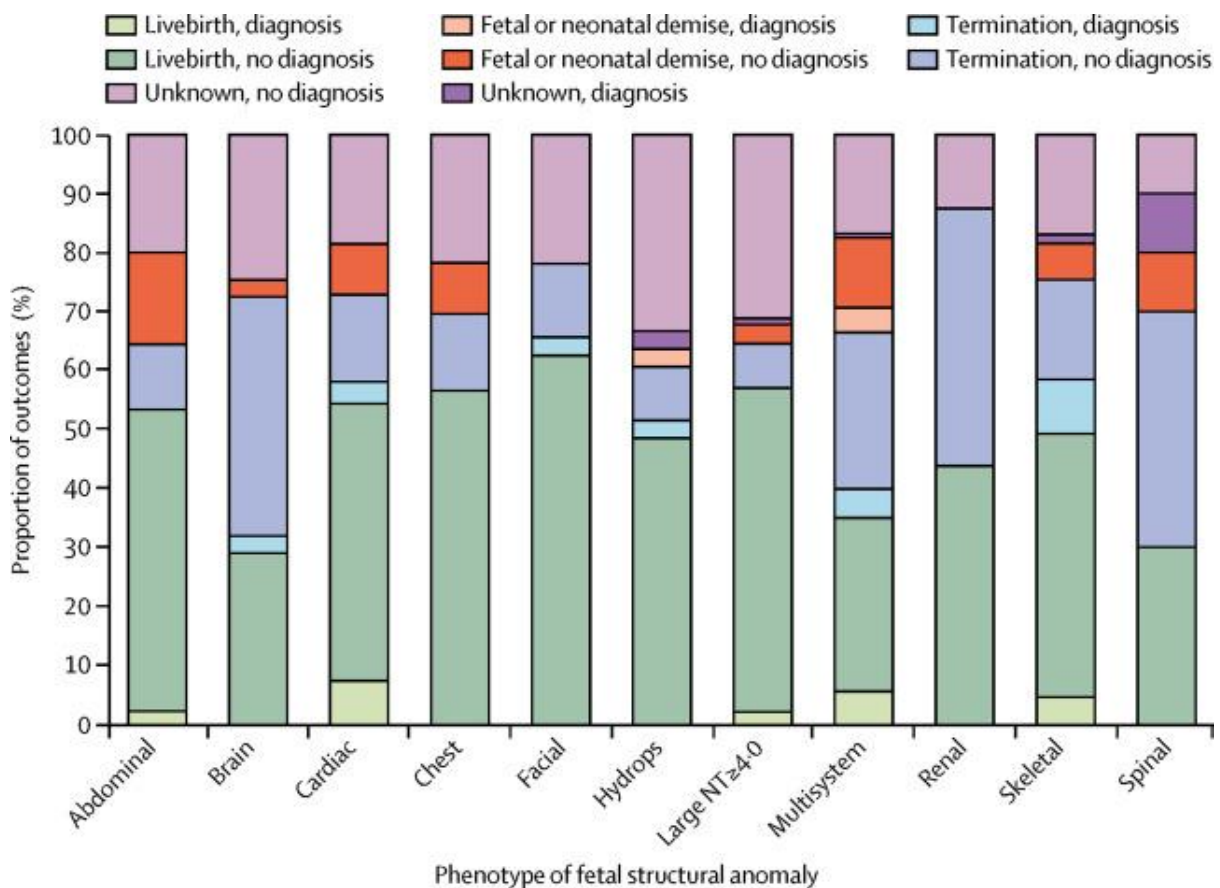


Fig Pregnancy outcomes associated with different fetal structural anomalies

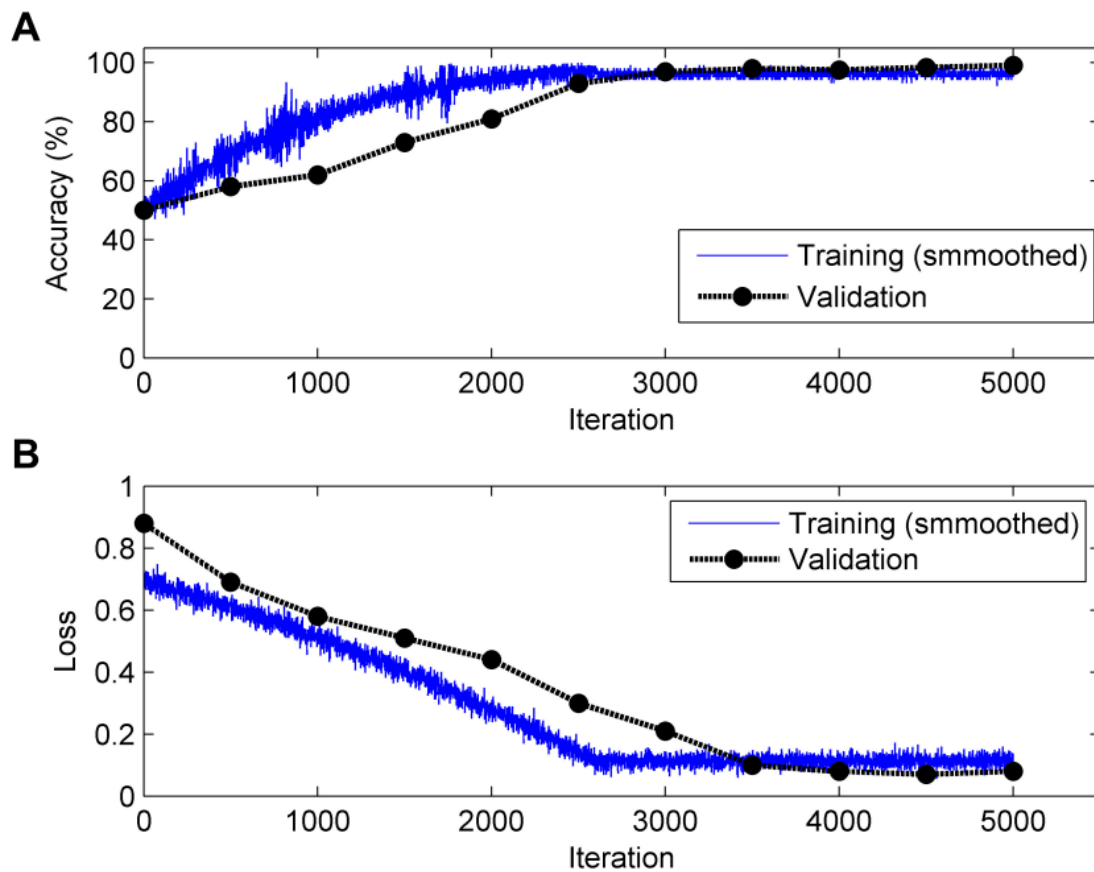


Fig: Training and Validation Process

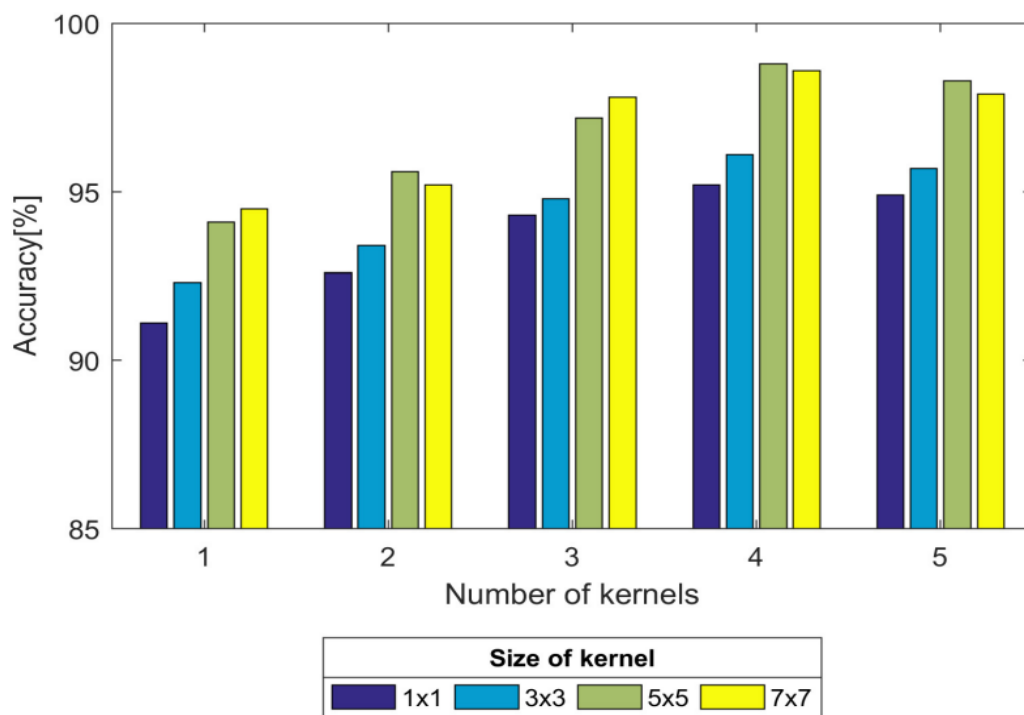


Fig : Accuracy For Difference Neural Network Models

4. Conclusion and Future Work:

Recently, CAD has played a significant role in predicting fetal development. The goal of this thesis work is to overcome the accuracy and sensitivity constraints of existing solutions in fetal growth and development, first, by developing an automated mass segmentation and classification system. In this study, the NormalShrink Homomorphic Filtering technique is used to compare various denoising filtering methods in the pre-processed stage. The results show that Normal Shrink Homomorphic Filtering Methods outperform spatial filters both subjectively and quantitatively, and that a genetic algorithm-based neuro-fuzzy classifier can discriminate between normal and abnormal mass.

The suggested methodology has obtained the best classification accuracy of 96 percent on the database of fetal pictures, compared to other current state-of-the-art classification methods such as ANN and GA based Neuro-Fuzzy. Method also showed to be the best when tested on both a benchmark database and a real database. As soon as prenatal ultrasound can discover congenital defects, there will be issues about what can and should be done. Everything involved in the prenatal diagnosis of syndromes is difficult due to the intricacy of the prenatal diagnosis of syndromes. This involves postnatal confirmation of the diagnosis, prognosis determination to assist parents in dealing with the unwell kid, and indication of the need for extensive, lifelong, and expensive multidisciplinary care of afflicted new-borns.

We also utilize the AUC of ROC as evaluation indicators to analyze the availability of feature data and classification accuracy more extensively. The following factors may have contributed to the encouraging results: The first is an improved segmentation method that can automatically locate and segment not only the mass but also with less reliance on the initial active contour and greater convergence capability; additionally, this segmentation method is robust to the interference of blurry areas and tissue and can precisely converge to the object. Last but not least, the genetic algorithm-based neuro-fuzzy classifier utilized in this study greatly enhances classification accuracy. The random forest was chosen for classification because of its unique properties: (1) it can deal with high-dimensional features without feature selection, (2) it can accommodate a large number of predictor variables, (3) it can reduce generalization error using unbiased estimation while creating random forests, and (4) it can run new data through previously generated forests to generate classifications. The suggested method may help the radiologist do a thorough examination of fetal growth and development in a short amount of time, improving the accuracy of fetal growth abnormalities diagnosis. Validation using a classification learner is the last stage, and SVM is the best validated strategy in terms of ROC curve 0.9998 area covered and Confusion Matrix accuracy rate of 97 percent.

References

- [1]. Maraci MA, Bridge CP, Napolitano R, Papageorgiou A, Noble JA. A framework for analysis of linear ultrasound videos to detect fetal presentation and heart beat. *J Med Image Anal* 2017; 37:22–36.
- [2]. Ali Liaqat. Intelligent image processing techniques for cancer progression detection, recognition and prediction in the human liver. *IEEE Symposium*;2014. <https://doi.org/10.1109/CICARE.2014.7007830>.
- [3]. Kalyan Karthik. Artificial neural network application in the diagnosis of disease conditions with liver ultrasound images. *Adv Bioinf* 2014.

- [4]. Persson M, Cnattingius S, Villamor E, et al. Risk of major congenital malformations in relation to maternal overweight and obesity severity: cohort study of 1.2 million singletons. *BMJ* 2017; 357: j2563.
- [5]. Wapner RJ, Martin CL, Levy B, et al. Chromosomal microarray versus karyo typing for prenatal diagnosis. *N Engl J Med* 2012; 367: 2175–84.
- [6]. Robson SC, Chitty LS, Morris S, et al. Evaluation of array comparative genomic hybridisation in prenatal diagnosis of fetal anomalies: a multicentre cohort study with cost analysis and assessment of patient, health professional and commissioner preferences for array comparative genomic hybridisation. *Effic Mech Eval* 2017; 4: 1–104.
- [7]. Hillman SC, Pretlove S, Coomarasamy A, et al. Additional information from array comparative genomic hybridization technology over conventional karyotyping in prenatal diagnosis: a systematic review and meta-analysis. *Ultrasound ObstetGynecol*2011; 37: 6–14.
- [8]. Talkowski ME, Ordulu Z, Pillalamarri V, et al. Clinical diagnosis by whole-genome sequencing of a prenatal sample. *N Engl J Med* 2012;367: 2226–32.
- [9]. Yang Y, Muzny DM, Xia F, et al. Molecular findings among patients referred for clinical whole-exome sequencing. *JAMA* 2014;312: 1870–79.
- [10]. Carss KJ, Hillman SC, Parthiban V, et al. Exome sequencing improves genetic diagnosis of structural fetal abnormalities revealed by ultrasound. *Hum Mol Genet* 2014; 23: 3269–77.
- [11]. Drury S, Williams H, Trump N, et al. Exome sequencing for prenatal diagnosis of fetuses with sonographic abnormalities. *Prenat Diagn* 2015; 35: 1010–17.
- [12]. Best S, Wou K, Vora N, Van der Veyver IB, Wapner R, Chitty LS. Promises, pitfalls and practicalities of prenatal whole exome sequencing. *Prenat Diagn* 2018; 38: 10–19.
- [13]. Chandler N, Best S, Hayward J, et al. Rapid prenatal diagnosis using targeted exome sequencing: a cohort study to assess feasibility and potential impact on prenatal counselling and pregnancy management. *Genet Med* 2018; published online March 29. DOI:10.1038/gim.2018.30.
- [14]. Calzolari E, Barisic I, Loane M, et al. Epidemiology of multiple congenital anomalies in Europe: a EUROCAT population-based registry study. *Birth Defects Res A Clin Mol Teratol.* 2014;100(4):270-276.2.
- [15]. Karim JN, Roberts NW, Salomon LJ, Papageorghiou AT. Systematic review of first-trimester ultrasound screening for detection of fetal structural anomalies and factors that affect screening performance. *Ultrasound Obstet Gynecol.* 2017;50(4):429-441.3.
- [16]. Skreden M, Skari H, Malt UF, et al. Long-term parental psychological distress among parents of children with a malformation—a prospective longitudinal study. *Am J Med Genet A.* 2010;152a(9):2193-2202.4.
- [17]. Boyle B, Addor MC, Arriola L, et al. Estimating global burden of disease due to congenital anomaly: an analysis of European data. *Arch Dis Child Fetal Neonatal Ed.* 2018;103(1): F22-f8.5.
- [18]. Wapner RJ, Martin CL, Levy B, et al. Chromosomal microarray vs karyo typing for prenatal diagnosis. *N Engl J Med.* 2012;367(23):2175-2184.6.
- [19]. Callaway JL, Shaffer LG, Chitty LS, Rosenfeld JA, Crolla JA. The clinical utility of microarray technologies applied to prenatal cytogenetics in the presence of a normal conventional karyotype: a review of the literature. *Prenat Diagn.* 2013;33(12):1119-1123.7.

- [20]. Neveling K, Feenstra I, Gilissen C, et al. A post-hoc comparison of the utility of sanger sequencing and exome sequencing for the diagnosis of heterogeneous diseases. *Hum Mutat.* 2013;34(12):1721-1726.8.
- [21]. Yang Y, Muzny DM, Reid JG, et al. Clinical whole-exome sequencing for the diagnosis of mendelian disorders. *N Engl J Med.* 2013;369(16):1502-1511.9.
- [22]. Stark Z, Tan TY, Chong B, et al. A prospective evaluation of whole-exome sequencing as a first-tier molecular test in infants with suspected monogenic disorders. *Genet Med.* 2016;18(11):1090-1096.10.
- [23]. Willig LK, Petrikin JE, Smith LD, et al. Whole-genome sequencing for identification of Mendelian disorders in critically ill infants: a retro-spective analysis of diagnostic and clinical findings. *Lancet Respir Med.* 2015;3(5):377-387.11.
- [24]. van Diemen CC, Kerstjens-Frederikse WS, Bergman KA, et al. Rapid targeted genomics in critically ill newborns. *Pediatrics.* 2017;140(4): e20162854.12.
- [25]. Meng L, Pammi M, Saronwala A, et al. Use of exome sequencing for infants in intensive care units: ascertainment of severe single-gene disorders and effect on medical management. *JAMA Pediatr.* 2017;171(12):e173438.13.
- [26]. Kingsmore SF, Cakici JA, Clark MM, et al. A randomized, controlled trial of the analytic and diagnostic performance of singleton and trio, rapid genome and exome sequencing in ill infants. *Am J Hum Genet.* 2019;105(4):719-733.14.
- [27]. Chandler N, Best S, Hayward J, et al. Rapid prenatal diagnosis using targeted exome sequencing: a cohort study to assess feasibility and potential impact on prenatal counseling and pregnancy management. *Genet Med.* 2018;20:1430-1437.15.
- [28]. Rydberg C, Tunon K. Detection of fetal abnormalities by second-trimester ultrasound screening in a non-selected population. *Acta Obstet Gynecol Scand.* 2017;96(2):176-182.16.
- [29]. Edwards L, Hui L. First and second trimester screening for fetal structural anomalies. *Semin Fetal Neonatal Med.* 2018;23(2):102-111.17.
- [30]. de Koning MA, Haak MC, Adama van Scheltema PN, et al. From diagnostic yield to clinical impact: a pilot study on the implementation of prenatal exome sequencing in routine care. *Genet Med.* 2019;21:2303-2310.18.
- [31]. Pangalos C, Hagnefelt B, Lilakos K, Konialis C. First applications of a targeted exome sequencing approach in fetuses with ultrasound abnormalities reveals an important fraction of cases with associated gene defects. *PeerJ.* 2016; 4: e1955.19.
- [32]. Normand EA, Braxton A, Nassef S, et al. Clinical exome sequencing for fetuses with ultrasound abnormalities and a suspected mendelian disorder. *Genome Med.* 2018;10(1):74.20.
- [33]. Daum H, Meiner V, Elpeleg O, Harel T. Fetal exome sequencing: yield and limitations in a tertiary referral center. *Ultrasound Obstet Gynecol.* 2019;53(1):80-86.982DEDENET AL.
- [34]. EUROCAT. Prenatal screening & diagnosis. Prenatal detection (pd) rates. Available at: [http://www.eurocatnetwork.eu/prenatalscreeninganddiagnosis/prenatal%20detection\(pd\)rates](http://www.eurocatnetwork.eu/prenatalscreeninganddiagnosis/prenatal%20detection(pd)rates). (Accessed on October 11, 2016).
- [35]. Kurjak A, Pooh RK, Merce LT, Carrera JM, Salihagic-Kadic A, Andonotopo W. Structural and functional early human development assessed by three-dimensional and four-dimensional sonography. *Fertil Steril.* 2005; 84:1285–99.
- [36]. Pooh RK. A New Field of “Fetal Sono-ophthalmology” by 3D HDlive Silhouette and Flow. *Donald School J Ultrasound Obstet Gynecol.* 2015; 9:221–2.

- [37]. Kurjak A, Miskovic B, Stanojevic M, Amiel-Tison C, Ahmed B, Azumendi G, et al. New scoring system for fetal neurobehavior assessed by three- and four-dimensional sonography. *J Perinat Med.* 2008; 36:73–81.
- [38]. Kurjak A, Abo-Yaqoub S, Stanojevic M, Yigiter AB, Vasilj O, Lebit D, et al. The potential of 4D sonography in the assessment of fetal neurobehavior-multicentric study in high-risk pregnancies. *J Perinat Med.* 2010; 38:77–82.
- [39]. Stanojevic M, Antsaklis P, Kadic AS, Predojevic M, Vladareanu R, Vladareanu S, et al. Is kurjak antenatal neurodevelopmental test ready for routine clinical application? Bucharest consensus statement. *Donald School J Ultrasound Obstet Gynecol.* 2015; 9:260–5.
- [40]. Kurjak A, Barišić LS, Stanojević M, Kadić AS, Porović S. Are we ready to investigate cognitive function of fetal brain? The role of advanced four-dimensional sonography. *Donald School J Ultrasound Obstet Gynecol.* 2016; 10:116–24.
- [41]. Barišić LS, Kurjak A, Pooh RK, Delić T, Stanojević M, Porović S. Antenatal detection of fetal syndromes by ultrasound: from a single piece to a complete puzzle. *Donald School J Ultrasound Obstet Gynecol.* 2016; 10:63–77.
- [42]. Pooh RK, Kurjak A. Novel application of three-dimensional HD live imaging in prenatal diagnosis from the first trimester. *JPerinat Med.* 2015; 43:147–58.
- [43]. Rawat Vidhi, Jain Alok, Shrimali Vibhakar, Rawat Abhishek. Automatic detection of fetal abnormality using head and abdominal circumference. *Int Conf Comput Collective Intell* 2016:25–534. https://doi.org/10.1007/978-3-319-45246-3_50.
- [44]. Subramanian Kalpathi R, Lawrence Dina M, Taghi Mostafavi M. Interactive segmentation and analysis of fetal ultrasound images.
- [45]. Pradeep Kumar BP, Prathap C, Dharshith CN. An automatic approach for segmentation of ultrasound liver images. *Int J Emerg Technol Adv Eng* 2013.
- [46]. Hiremath PS, Tegnoor Jyothi R. Fuzzy inference system for follicle detection in ultrasound images of ovaries. *Soft Comput* 2014;18(7):1353–62.
- [47]. Harsh Vardhan M et al. GLCM architecture for image extraction. *Int J Adv Res Electron Commun Eng (IJARECE)* 2014;3(1).
- [48]. Kalyan Karthik. Application of artificial neural networks towards the determination of presence of disease conditions in ultrasound images of kidney. *Int J Comput Eng Technol* 2013;4(5):232–43.
- [49]. Singh Mandeep, Singh Sukhwinder, Gupta Savita. An information fusion-based method for liver classification using texture analysis of ultrasound images. *Inf Fusion* 2014; 19:91–6.