# REVIEW

# Computational Intelligence in Early Diabetes Diagnosis: A Review

Shankaracharya[1], Devang Odedra[1], Subir Samanta[2], and Ambarish S. Vidyarthi[1]

[1] Department of Biotechnology, Birla Institute of Technology, Mesra, Ranchi 835215, India. [2] Department of Pharmaceutical Sciences, Birla Institute of Technology, Mesra, Ranchi 835215, India. Address correspondence to: Shankaracharya, e-mail: shankaracharya@bitmesra.ac.in

## ■ Abstract

The development of an effective diabetes diagnosis system by taking advantage of computational intelligence is regarded as a primary goal nowadays. Many approaches based on artificial network and machine learning algorithms have been developed and tested against diabetes datasets, which were mostly related to individuals of Pima Indian origin. Yet, despite high accuracies of up to 99% in predicting the correct diabetes diagnosis, none of these approaches have reached clinical application so far. One reason for this failure may be that diabetologists or clinical investigators are sparsely informed about, or trained in the use of, computational diagnosis tools. Therefore, this article aims at sketching out an outline of the wide range of options, recent developments, and potentials in machine learning algorithms as diabetes diagnosis tools. One focus is on supervised and unsupervised methods, which have made significant impacts in the detection and diagnosis of diabetes at primary and advanced stages. Particular attention is paid to algorithms that show promise in improving diabetes diagnosis. A key advance has been the development of a more in-depth understanding and theoretical analysis of critical issues related to algorithmic construction and learning theory. These include trade-offs for maximizing generalization performance, use of physically realistic constraints, and incorporation of prior knowledge and uncertainty. The review presents and explains the most accurate algorithms, and discusses advantages and pitfalls of methodologies. This should provide a good resource for researchers from all backgrounds interested in computational intelligence-based diabetes diagnosis methods, and allows them to extend their knowledge into this kind of research.

**Keywords**: diabetes diagnosis · computational · algorithm · artificial neural network · learning · logistic regression

## Introduction

Diabetes has been recognized as a continuing health challenge for the twenty-first century, both in developed and developing countries. It is understood that diabetes prevalence is increased because of modern lifestyles, urbanization, and economic development [1]. It is a global problem with devastating human, social, and economic impact, affecting around 300 million people worldwide [2].

Type 2 diabetes is a chronic disease that occurs either when the pancreas does not produce enough insulin, or when the body cannot effectively use the insulin it produces. It is frequently asymptomatic [3]. Although detection is improving, the delay from disease onset to diagnosis may exceed 10 years [4]. To diagnose diabetes, a physician has to analyze many factors. Undoubtedly, the evaluations of data obtained from patients and expert decisions are critical for diagnosis. However, factors such as lack of experience by the ex-

perts, or their fatigue, may lead to erroneous diagnosis. Early intervention with lifestyle modifications or pharmacotherapy has been shown to effectively delay or prevent type 2 diabetes and its complications in adults [5].

---

**Abbreviations**:

ADAP - adaptive learning routine
ANFIS - artificial neuro-fuzzy inference system
ANN - artificial neural network
ARTMAP - adaptive resonance theory mapping
ARTMAP-IC - adaptive resonance theory mapping instance counting
BPNN - back-propagation neural network
CART - classification and regression trees
CART-DB - classification and regression trees distribution-based
ESOM - evolving self-organizing maps
FIS - fuzzy inference system
GCS - growing cell structure
GDA - generalized discriminant analysis
GNG - growing neural gas
GRG2 - generalized reduced gradient 2
GRNN - general regression neural network
k-NN - k-nearest neighbor
LDA - linear discriminant analysis
LM - Levenberg-Marquardt
LS-SVM - least square support vector machine
LVQ - learning vector quantization
ME - mixture of experts
MEA - multimodal evolutionary algorithm
MFNNCA - modified feed forward neural network constructive algorithm
MKS - multiple knot spline
MLP - multi-layer perceptron
MLPNN - multi-layer perceptron neural network
MLNN - multilayer neural networks
MME - modified mixture of experts
NFIS - neuro-fuzzy inference system
NG - neural gas
NHANES - National Health and Nutrition Examination Survey
PC - principal components
PCA - principal component analysis
PID - Pima Indian diabetes dataset
PNN - probabilistic neural network
RBF - radial basis function
SOM - self-organizing map
SSVM - smooth support vector machines
SVM - support vector machine
UCI - University of California, Irvine

---

For prevention of type 2 diabetes, a comprehensive guideline was issued specifying lifestyle changes [6]. Various strategies have also been put forward to reduce diabetes risk [7]. Naturally, prevention is preferable, but current treatment methods are not yet fully adequate to reach this goal. Hence, there is a growing need for early detection of diabetes. To address this need, and to provide more detailed and rapid analysis of medical data,

risk assessment tools and their various algorithms have been widely investigated.

For early detection of diabetes, various risk scores have been devised. A detailed survey of these tools with their specificity and sensitivity has been provided by Schwarz *et al.* in which the authors found the Finnish Diabetes Risk Score as the most convenient tool for early diagnosis of diabetes [8]. However, as this method involves human intervention in deciding criteria and score, it may by be exposed to the human error. Therefore, machine learning and statistical pattern recognition has been the subject of tremendous interest in the biomedical community as these approaches offer promise for improving the sensitivity and/or specificity of detection and diagnosis of disease. At the same time, these approaches reduce the potential for human error in the decision making process [9]. In particular, further development of methods that explicitly incorporate prior knowledge and uncertainty into the decision-making process would be very important for diabetes detection. Extensive studies by many researchers have demonstrated higher performance and accuracy in predicting clinical outcomes of diabetes diagnosis using neural network strategies (Table 1). Advantages and pitfalls of using various algorithms in diabetes prediction are listed in Table 2.

## Datasets for diabetes diagnosis

Significant work has been reported on Pima Indian diabetes datasets (PID). These studies applied different methods to the given problem, and achieved high classification accuracies using the dataset taken from the University of California, Irvine (UCI) machine learning repository [10]. This database provides a well validated data resource to explore the prediction of diabetes. The eight variables in the dataset include:

- number of times pregnant,
- plasma glucose concentration at 2 hour in an oral glucose tolerance test,
- diastolic blood pressure (mmHg),
- triceps skin fold thickness (mm),
- 2-h serum insulin (IU/ml),
- body mass index (weight in kg/height in m),
- diabetes pedigree function, and
- age (years).

While PID is one of the mostly used datasets for prediction of type 2 diabetes, some researchers prefer to investigate diagnosis using data from hospitals, and to incorporate their own parameters

**Table 1.** Artificial intelligence approaches for early diabetes detection

| Algorithm | Dataset | Accuracy (%) | Specificity (%) | Sensitivity (%) | Reference |
|---|---|---|---|---|---|
| MFNNCA | PID | 80.07 | 84.38 | 74.00 | Kamaruzzaman et al. [51] |
| GRG2 | PID | 81.25 | - | - | Shanker et al. [15] |
| ANFIS | PID | 98.14 | 98.58 | 96.97 | Ubeyli [52] |
| GRNN | PID | 80.21 | - | - | Kayaer, Yildirim [38] |
| MLP | PID | 77.08 | - | - | Kayaer, Yildirim [38] |
| RBF | PID | 68.23 | - | - | Kayaer, Yildirim [38] |
| ARTMAP-IC | PID | 81.00 | - | - | Carpenter, Markuzon [47] |
| MEA | PID | 80.07 | - | - | Stoean et al. [53] |
| ESOM | PID | 78.40 | - | - | Deng, Kasabov [23] |
| GNG | PID | 74.60 | - | - | Deng, Kasabov [23] |
| GCS | PID | 73.80 | - | - | Deng, Kasabov [23] |
| k-NN | PID | 77.00 | - | - | Kordos et al. [16] |
| k-NN | PID | 71.90 | - | - | Ster, Dobnikar [17] |
| CART | PID | 72.80 | - | - | Ster, Dobnikar [17] |
| MLP | PID | 75.20 | - | - | Ster, Dobnikar [17] |
| LVQ | PID | 75.80 | - | - | Ster, Dobnikar [17] |
| LDA | PID | 77.50 | - | - | Ster, Dobnikar [17] |
| CART-DB | PID | 74.40 | - | - | Shang, Breiman [54] |
| SVM | Questionnaire | 94.00 | 94.00 | 93.00 | Barakat et al. [25] |
| SSVM | PID | 76.73 | - | - | Purnami et al. [27] |
| MKS-SSVM | PID | 93.20 | - | - | Purnami et al. [27] |
| GDA and LS-SVM | PID | 78.21 | - | - | Polat et al. [44] |
| PCA-ANFIS | PID | 89.47 | - | - | Polat, Gunes [55] |
| LDA-ANFIS | PID | 84.61 | 85.18 | 83.33 | Dogantekin et al. [44] |
| Naive Bayes | PID | 74.50 | - | - | Friedman [56] |
| Semi-naive Bayes | PID | 76.00 | - | - | Friedman [56] |
| C4.5 | PID | 76.00 | - | - | Friedman [56] |
| MLPNN | PID | 91.53 | 91.19 | 92.42 | Ubeyli [49] |
| ME | PID | 97.93 | 98.01 | 97.73 | Ubeyli [49] |
| MME | PID | 99.17 | 99.43 | 98.48 | Ubeyli [49] |

**Legend**: PID: Pima Indian dataset. MFNNCA: modified feed forward neural network constructive algorithm. GRG2: generalized reduced gradient 2. ANFIS: adaptive neuro-fuzzy inference system. GRNN: general regression neural network. MLP: multi-layer perceptron. RBF: radial basis function. ARTMAP-IC: adaptive resonance theory mapping instance counting. MEA: multimodal evolutionary algorithm. ESOM: evolving self-organizing maps. GNG: growing neural gas. GCS: growing cell structure. k-NN: k-nearest-neighbor. CART: classification and regression trees. LVQ: learning vector quantization. LDA: linear discriminant analysis. CART-DB: classification and regression trees distribution-based. SVM: support vector machine. SSVM: smooth support vector machine. MKS-SSVM: multiple knot spline smooth support vector machine. GDA: generalized discriminant analysis. LS-SVM: least square support vector machine. PCA-ANFIS: principal component analysis and adaptive neuro-fuzzy inference system. LDA-ANFIS: linear discriminant analysis and adaptive network based fuzzy inference system. C4.5: sample class 4.5 algorithm. MLPNN: multi-layer perceptron neural network. ME: mixture of experts. MME: modified mixture of experts.

of interest. Kazemnejad *et al.* used the Tehran Lipid and Glucose Study dataset which consists of variables like age, body mass index, waist-to-hip ratio, gender, history of hyperlipidemia, and history of hypertension [11]. In another study conducted by Dey *et al.* on data of 530 patients from Sikkim Manipal Institute of Medical Sciences, risk factors such as random blood sugar test results, fasting blood sugar test results, post plasma blood

sugar tests, age, sex, and occupation were taken into account [12].

The third National Health and Nutrition Examination Survey (NHANES III, http://www.cdc.gov/diabetes/) dataset resulted from a survey conducted on a US population. The eighteen variables identified as important for diabetes risk prediction include body mass index, height, weight, waist circumference, waist-to-hip ratio, age, sex,

race/ethnicity, taking blood pressure medication, taking cholesterol medication, gestational diabetes, high blood pressure, high cholesterol, history of diabetes (any blood relative), history of diabetes (parent or sibling), history of diabetes (parent), history of diabetes (sibling), and exercise [13].

# Data analysis through logistic regression

Logistic regression can be applied when the data consist of a binary response and a set of explanatory variables [14]. At first, the maximum likelihood estimates for the parameters of the logistic regression model are estimated using an iteratively reweighted least squares algorithm. Then, it is possible to calculate the predicted probability of an individual having diabetes by using the following logistic function:

$$\theta = \frac{1}{1 + e^{-\left(\beta_0 + \beta_1 x_1 + \beta_2 x_2 + \dots + \beta_n x_n\right)}}$$

Here $X$ is a vector of variables and $\beta$ is the regression coefficient estimated by using maximum likelihood methods. Shanker applied logistic regression on eight variables in PID and obtained a significant accuracy of 79.17% [15]. Statistically least significant (at 0.05 level) variables were deleted sequentially in the training sample. Logistic regression with the remaining four statistically significant parameters, e.g. number of times pregnant, glucose tolerance test, body mass index, and diabetes pedigree function, resulted in an overall classification accuracy of 80.21%. Heikes *et al.* have developed a diabetes risk calculator tool based on logistic regression function to identify people at high risk of diabetes [13]. It was built upon NHANES III dataset with a sensitivity of 75%.

## Clustering techniques

Most quality prediction models are based on clustering techniques that make use of k-means, mixture-of-Gaussians, self-organizing map (SOM) and neural gas (NG) for diagnosis. According to

**Table 2.** Advantages and disadvantages of algorithms commonly used in diabetes prediction

| Algorithm | Advatages | Disadvantages |
|---|---|---|
| Back propagation | Better error minimization | Slow convergence rate |
| LM | Fast convergence rate | Memorization effect on over-training |
| SVM | Guaranteed global minimum | No specific rule to choose a kernel that will give better classification |
| ANFIS | Fast convergence rate | Low interpretability of learned information, computationally expensive |
| RBF | Uses small numbers of locally tuned units and is adaptive in nature | Sensitive to dimensionality of data |
| ARTMAP-IC | Fast convergence rate | Tends to be conservative which reduces sensitivity |
| SOM | Little computational and memory requirements | Topology mismatch leads to poor classification |
| ESOM | Shorter learning process than SOM | Poor adaptability to input data |
| GNG | Can adaptively determine the number of connections | Poor response to changing inputs |
| k-NN | Good choice when there is no prior knowledge of data distribution | Requires rigorous tuning to optimally fit the real world data |
| LVQ | Little computational and memory requirements | Less accurate with high dimensional data |
| LDA | Works best when class has Gaussian density | Less accurate with small sample size |
| ME | Requires only small number of connections in neural network | Learns only static input-output mappings (i.e. no feedback) |
| MME | Requires only small number of connections in neural network. Faster than ME | Learns only static input-output mappings |

**Legend**: SVM: support vector machine. ANFIS: adaptive neuro-fuzzy inference system. RBF: radial basis function. ARTMAP-IC: adaptive resonance theory mapping instance counting. SOM: self-organizing maps. ESOM: evolving self-organizing maps. GNG: growing neural gas. k-NN: k-nearest-neighbor. LVQ: learning vector quantization. LDA: linear discriminant analysis. ME: mixture of experts. MME: modified mixture of experts.

the k-nearest neighbor (k-NN) algorithm, a new input pattern **x** is assigned to the class voted by the majority of its k-nearest training patterns [16]. The weight change in k-NN is given by:

$$\Delta W_j = f(x) = \begin{cases} \gamma(X - W_j), & \text{if } j = i(X) \\ 0, & \text{otherwise} \end{cases}$$

where $\gamma$ is the learning rate and i(x) is the winning node. While the accuracy of k-NN on diabetes detection problem ranges between 71-78% [16, 17], a more sensitive performance with accuracy of 92.38% was achieved with a hybrid model of k-NN and C4.5 algorithms [18, 19].

SOM is a sheet-like artificial neural network (ANN). Cells of this ANN become specifically tuned to input patterns [20]. In order to overcome the topology mismatches that occur with the original SOM algorithm, and to achieve an optimal use of the neurons, the geometry of the lattice has to match with the data manifold. For this purpose, several so-called growing (incremental) SOM algorithms have been developed. The growing neural gas (GNG) algorithms start with two randomly placed, connected neurons [21]. After a fixed number of time steps, the neuron $i$ with the largest accumulated error is determined, and a new neuron inserted between $i$ and one of its neighbors. It does not require predetermination of the neuron quantity or topology of structure to be used. It starts with a minimal neuron structure that is incremented during training until it reaches a maximum number limit for clusters defined by the user.

The growing cell structure (GCS) algorithm assumes a fixed dimensionality for the lattice [22]. It is well suited for generating a dimensionality-reducing mapping from the input space to the lattice space. Deng and Kasabov applied GNG and GCS algorithms to the diabetes diagnosis problem, and reported accuracies of 74.6% and 73.8%, respectively [23]. Both GNG and GCS need to calculate local resources for prototypes, which introduces extra computational effort and reduces their efficiency. Deng and Kasabov proposed the evolving self-organizing maps (ESOM) network structure, which is similar to that of GNG [21]. When applied to diabetes diagnosis, they obtained 78.4% classification accuracy using ESOM.

## Support vector machine (SVM)

Support vector machine (SVM) operates by finding a linear hyperplane that separates the positive and negative examples with a maximum interclass distance [24]. We can define $z_i$ as an indicator variable which specifies whether a data vector $\mathbf{x}_i$ is in class diabetics or non-diabetics (e.g., $z_i = -1$ if $\mathbf{x}_i$ is in the diabetic class and $z_i = 1$ if $\mathbf{x}_i$ is in the non-diabetic class). The distance of a hyperplane $\mathbf{w}$ to a (transformed) data vector $\mathbf{y}$ is defined as $|f(\mathbf{y})|/||\mathbf{w}||$. Together with the fact that the separating hyperplane ensures $z_i f(\mathbf{y}_i) \geq 1$ for all $n$ data vectors $i$, we can express the condition on the margin $m$ as:

$$\frac{z_i f(\mathbf{y})}{||\mathbf{w}||} \geq m, \text{ where } i = 1,...,n$$

The goal of SVM training is to find the weight vector **w** that maximizes the margin $m$. Barakat *et al.* employed SVM to process the inputs, and extracted the rules using an electic approach [25]. This approach was then used to predict the diagnosis of diabetes using a questionnaire based on demographic, historic, and anthropometric measures. The authors achieved a prediction accuracy of 94%.

A cascade learning system based on generalized discriminant analysis (GDA) and least square support vector machine (LS-SVM) has been proposed for early diagnosis of Pima Indian diabetes disease [26]. The accuracy reported in this study was 78.21% with 10-fold cross-validation. Purnami *et al.* applied smooth support vector machines (SSVM) to the diabetes detection problem [27]. SSVM, developed by Lee *et al.*, is an extension to SVM in which smoothing function is applied to solve the problem [28]. With SSVM, the investigators achieved a 76.73% accuracy. To improve efficiency, they proposed a new multiple knot spline (MKS) smoothing function for SSVM. Replacing the default-plus function of SSVM by MKS, they enhanced the automated diagnosis performance of SSVM with an accuracy of 93.2%.

## Neural networks

### Multi-layer neural networks

Multilayer neural networks (MLNN) are composed of one or more hidden layers between input and output (Figure 1) [29]. In the training phase, the training data is fed through the input layer. The data is propagated from the hidden layer to the output layer (Figure 2), which is called forward pass. During this phase, each node in the hidden layer gets input from all the input layer
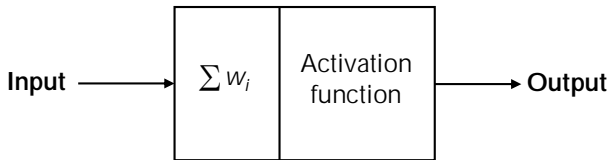
**Figure 1.** Architecture of a single neuron.

nodes, which are then multiplied by the randomly assigned weights before summing up. Similarly, the output layer node receives inputs from all nodes of the hidden layer, which are then multiplied by the randomly assigned weights and summed up. This forms the output of the output layer.

The input to each hidden layer is calculated by:

$$y = \sum w_i * \text{input value}$$

where $w_i$ is the weight for neuron $i$. The output of the hidden layer is calculated by using an activation function. The activation function acts as a squashing function, such that the output of a neuron in a neural network is between certain values (usually 0 and 1 for sigmoid, or -1 and 1 for hyperbolic tangent). Common activation functions used in diabetes diagnosis are the sigmoid (a) and hyperbolic tangent (b) function:

$$\text{a)} \ \ f(x) = \frac{1}{1 + e^{-x}} \qquad \text{b)} \ \ f(x) = \frac{e^x - e^{-x}}{e^x + e^{-x}}$$

with sigmoid range = [0,1], and hyperbolic range = [1,-1]. Error rates are calculated as follows:

$$\text{Error} = f(x) * \left[1 - f(x)\right] * \left[\text{target value} - f(x)\right]$$
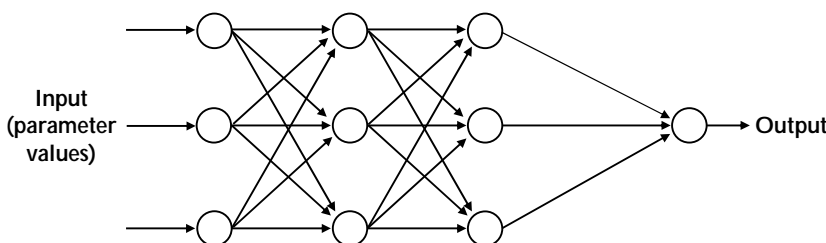


**Figure 2.** Multi-layer neural network with 3 neuron layers.

## Back-propagation neural networks

The back-propagation neural network (BPNN) algorithm is widely recognized as a powerful tool for training of the MLNN. In this algorithm, errors are back-propagated to the hidden layers, weights are reassigned, and the process continues until the error rate is a minimum. The new weights are calculated based on the following equation:

$$\text{Weight (new)} = \text{weight} + \eta * \text{error} * f(x)$$

where $\eta$ is the learning rate. However, since it applies the steepest descent method to update the weights, it suffers from a slow convergence rate, and often yields suboptimal solutions [30, 31]. Jaafar *et al.* used the back propagation neural network algorithm for diagnosing diabetes [32]. The inputs to the system were glucose tolerance test, diastolic blood pressure, triceps skin fold thickness, serum insulin, body mass index, diabetes pedigree function, number of times pregnant, and age. BPNN was used to predict the glucose level [33], and also to train and test its performance using diabetes patients [12].

Although the BPNN algorithm is widely used, one major drawback is that it requires a complete set of input data. However, most diabetes datasets are often incomplete in the one respect or another. Back propagation algorithm cannot interpret the missing values (if any) which may prevent the identification of factors leading to rare outputs. To overcome this problem, Jayalakshmi and Santhakumaran proposed a new approach to deal with the missing values [34]. They achieved an accuracy of 99.9% by replacing the missing values with its mean, and then normalizing the data with a principal component analysis (PCA) technique [35]. PCA is an extraction method aimed at describing the data variance by constructing a set of new orthogonal features called principal components (PCs). The PCs are a linear combination of the data variables that are mutually orthogonal. Every new PC describes a part of the data variance not explained by components used previously. Due to this fact, a few first PCs are usually enough to represent the data variance well.

It was reported that the Levenberg-Marquardt (LM) al-

gorithm [36] provides generally faster convergence and better estimation results than other training algorithms [37]. However, this method can cause a memorization effect when overtraining occurs. If a neural network starts to memorize the training set, its generalization starts to decrease, and its performance may not be improved for untrained test sets. Kayaer and Yildirim used the LM algorithm on a Pima Indian dataset, and achieved an accuracy of 77.08% [38], which was lower than other algorithms. Temurtas *et al.* trained the neural network optimally with a probabilistic neural network (PNN) along with a LM algorithm [39, 40]. They achieved an 82.37% accuracy with this approach.

### Radial basis function (RBF)

In neural networks, radial basis functions (RBFs) are used as a replacement for the sigmoidal hidden layer transfer function in multi-layer perceptrons (MLP) [41]. The only parameters adjusted in the learning process are the linear mapping from the hidden layer to the output layer. Hence, RBF networks have the advantage of not suffering from local minima.

RBF shows good performance in regression applications where the input space dimension is relatively small. However, in prediction problems like diabetes diagnosis, only 68.23% efficiency has been reported, which is far less than other algorithms. RBF networks have the disadvantage of requiring good coverage of the input space by radial basis functions. Determination of RBF centers is heavily dependent on the distribution of the input data without reference to the prediction task.

### General regression neural network (GRNN)

The general regression neural network (GRNN) is related to the radial basis function network and is based on a standard statistical technique called Kernel regression [42]. It approximates any arbitrary function between input and output vectors, and draws the function estimate directly from the training data. It does not require an iterative training procedure, as in MLP. For an input estimator 'x', corresponding to diabetes risk factor variables, GRNN produces an output estimator 'y' which minimizes the estimation error. GRNN works on following formula:

$$E(y|x) = \int_{\infty}^{-\infty} yf(x, y)\, dy \Big/ \int_{\infty}^{-\infty} f(x, y)\, dy$$

where $E[y|x]$ is the expected value of output $y$, given the input vector $x$, and $f(x, y)$ the joint probability density function of $x$ and $y$.

GRNNs produce a real-valued prediction between 0 and 1. A cut-off value decides the criteria to identify positive prediction. The best result achieved by GRNN on PID is 80.21% using 0.5 as cut-off value for the decision [38].

### Neuro-fuzzy inference systems (NFIS)

A neuro-fuzzy network is a fuzzy inference system in an artificial neural network [43]. Depending on the fuzzy inference system (FIS) type, there are several layers that simulate the processes involved in a fuzzy inference like fuzzification, inference, aggregation, and defuzzification. Embedding a FIS in the general structure of an artificial neural network (ANN) has the benefit of using ANN training methods to find the parameters of a fuzzy system. Linear discriminant analysis (LDA) is used to separate the two types of feature variables in a given dataset [44]. Dogantekin *et al.* used LDA along with artificial neuro FIS (ANFIS) for the detection of diabetes [45]. In this method, LDA is used to separate feature variables between healthy and diabetes data. In the second phase, both the healthy and diabetes features obtained in the first phase are given to inputs of the ANFIS classifier. They achieved an 84.61% accuracy with this approach.

Smith *et al.* used the PID data set to evaluate the perceptron-like adaptive learning routine (ADAP), and achieved a prediction accuracy of 76% [46]. The performance of fuzzy adaptive resonance theory mapping (ARTMAP) on the same database was 66% [47]. ARTMAP is a supervised learning algorithm for input binary vectors. However, the ARTMAP algorithm required fewer rules and was comparatively faster. Carpenter and Markuzon have presented an instance counting algorithm (ARTMAP-IC) and obtained an 81% accuracy against the test set [47].

## Expert systems

In real world problems like diabetes detection, a simple classifier is too weak for accurate prediction. The use of expert systems and different artificial intelligence techniques for classification systems in medical diagnosis is increasing gradually. Mixture of experts and modified mixture of experts have been successfully implemented to the problem of diabetes diagnosis prediction.

## Mixture of experts

The new supervised learning algorithm called mixture of experts (ME) was proposed by Jacobs *et al.* [48]. This algorithm divides a learning task into appropriate subtasks, each of which can be solved by simple expert network. The global output of the ME system is derived as a convex combination of the outputs from a set of N experts, in which the overall predictive performance of the system is generally superior to any of the individual experts.
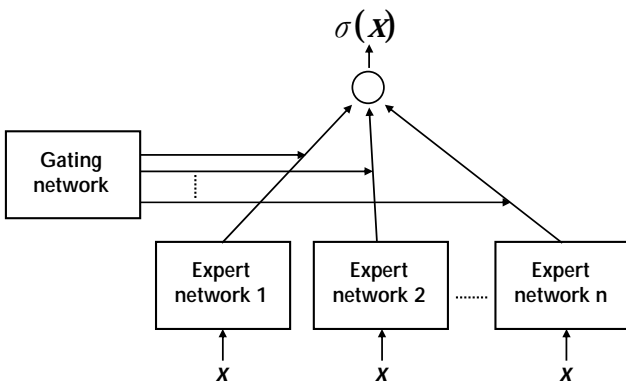


**Figure 3**. General architecture of mixture of experts.

ME architecture is composed of several expert networks and a gating network (Figure 3). The gating network produces a scalar output from a vector input $X$. The gating network operates on a generalized linear function where the output for $i^{th}$ input variable is given by:

$$g(x, v_i) = \frac{e^{\xi_i}}{\sum\limits_{k=1}^{n} e^{\xi_k}}$$

where $\xi_i = V_i^T x$, and $V_i$ is the weight vector. Each expert network produces an output vector for an input vector based on the following generalized linear equation:

$$\sigma_i(x) = f(W_i x)$$

where $W_i$ is a weight matrix. The final output of ME is the sum of multiplications of the outputs from gating and expert networks:

$$\sigma(x) = \sum_{k=1}^{n} g(x, v_k) \sigma_k(x)$$

Ubeyli presented an approach to test the performance of ME on PID with a classification accuracy of 97.93% [49], which was better than conventional MLNN. Moreover, the computational time required for classification using ME was comparatively small.

## Modified mixture of experts (MME)

Ubeyli [49] employed a new, fast, and effective modified mixture of experts (MME) approach proposed by Chen [50] to further improve the classification accuracy of ME.

The MME architecture is composed of an assembly of N expert networks and a gate-bank (Figure 4). For k different features, expert networks are divided into k groups, each comprising of N expert networks. Similarly, the gate-bank is composed of k gating networks. The resultant output of the gate-bank is a convex weighted sum of outputs produced by all the gating networks. Finally, the overall output of MME is obtained by linear combination of outputs of all N expert networks weighted by the output of the gate-bank.
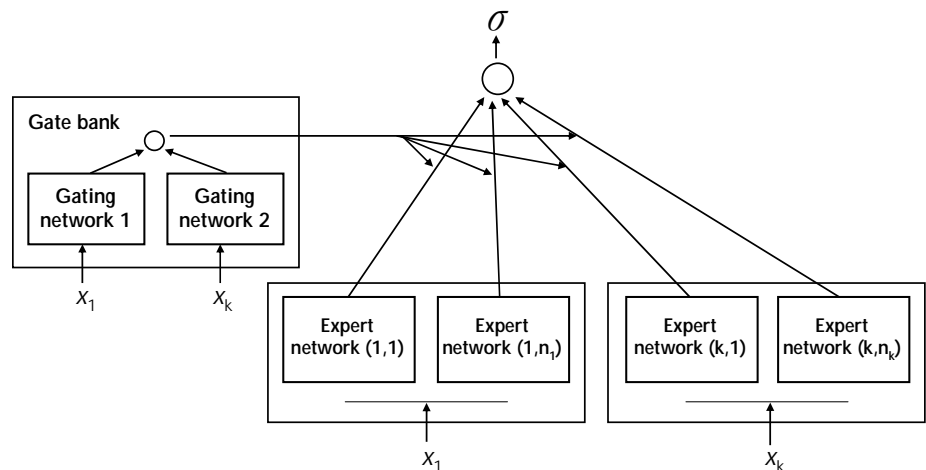


**Figure 4**. General architecture of modified mixture of experts.

Ubeyli applied the MME algorithm to the diabetes diagnosis problem and achieved an accuracy of 99.17% [49]. Apart from outperforming all other algorithms, the computational time required for classification was very small.

## Conclusions

Despite of the rapid development of theories for computational intelligence, application to diabetes diagnosis remains a challenge. This is due to specific problems of data use. These problems arise when statistical models of data are unknown or time-dependent, or when the parameters of the learning system need to be updated incrementally, while only a partial glimpse of incoming data is available. Based on the promising outcomes of studies applying computational algorithms to the problem of diabetes diagnosis, it is clear that a more sophisticated risk score could be developed. This would significantly decrease healthcare costs via early prediction and diagnosis of type 2 diabetes.

Some algorithms work better on the diabetes diagnosis problem than others. It will be important to compare outcomes further to find the most reliable algorithm for clinical application. Neural network methodology has outperformed classical statistical methods in cases where input variables are interrelated. Because clinical measurements are usually derived from multiple interrelated systems, it is evident that neural networks might be more accurate than classical methods in multivariate analysis of clinical data.

Trained models of diabetes risk factors should be incorporated into easy-to-use software solutions such that medical practitioners, who are not experts in artificial intelligence and computational techniques, may apply them easily. For this purpose, graphical user interface-enabled tools need to be developed by which medical practitioners can simply enter health profiles of their patients and receive an instant diabetes prediction with an acceptable degree of confidence. If the ANN-based prediction approach shows improved medical diagnosis, then it may become more widely accepted as a means to assist patient care in more hospitals and clinics.

Though the PID dataset provides a well validated data for predicting diabetes diagnosis, it is possible that models trained on such a dataset may not perform equally well on profiles of patients from other ethnic group. Therefore, it is recommended that models of choice must be trained on a dataset that closely represents patient profiles of medical practitioners within specific geographic regions.

**Disclosures (conflict of interests statement)**: The authors report no conflict of interests.

## ■ References

1. **Lieberman LS.** Dietary, evolutionary, and modernizing influences on the prevalence of type 2 diabetes. *Annu Rev Nutr* 2003. 23:345-377.
2. **Zhang Y, Dall T, Mann SE, Chen Y, Martin J, Moore V, Baldwin A, Reidel VA, Quick WW.** The economic costs of undiagnosed diabetes. *Popul Health Manag* 2009. 12(2):95-101.
3. **Jackson DM, Wills R, Davies J, Meadows K, Singh BM, Wise PH.** Public awareness of the symptoms of diabetes mellitus. *Diabet Med* 1991. 8:971-972.
4. **Harris MI, Klein R, Wellborn TA, Knuiman MW.** Onset of NIDDM occurs at least 4-7 yr before clinical diagnosis. *Diabetes Care* 1992. 15:815-819.
5. **Knowler WC, Barrett-Connor E, Fowler SE, Hamman RF, Lachin JM, Walker EA, Nathan DM.** Reduction in the incidence of type 2 diabetes with lifestyle intervention or metformin. *N Engl J Med* 2002. 346(6):393-403.
6. **Paulweber B, Valensi P, Lindström J Lalic NM, Greaves CJ, McKee M, Kissimova-Skarbek K, Liatis S, Cosson E, Szendroedi J, et al.** A European evidence-based guideline for the prevention of type 2 diabetes. *Horm Metab Res* 2010. 42(Suppl 1):S3-S36.
7. **Lindstrom J, Neumann A, Sheppard KE, Gilis-Januszewska A, Greaves CJ, Handke U, Pajunen P, Puhl S, Pölönen A, Rissanen A, et al.** Take action to prevent diabetes - the IMAGE toolkit for the prevention of type 2 diabetes in Europe. *Horm Metab Res* 2010. 42 (Suppl 1):S37-S55.
8. **Schwarz PE, Li J, Lindstorm J, Tuomilehto J.** Tools for predicting the risk of type 2 diabetes in daily practice. *Horm Metab Res* 2009. 41(2):86-97.
9. **Sajda P.** Machine learning for detection and diagnosis of disease. *Annu Rev Biomed Eng* 2006. 8:537-565.
10. **Frank A, Asuncion A.** UCI machine learning repository. Irvine, CA, University of California, School of Information and Computer Science, 2010.
11. **Kazemnejad A, Batvandi Z, Faradmal J.** Comparison of artificial neural network and binary logistic regression for determination of impaired glucose tolerance/diabetes. *East Mediterr Health J* 2010. 16(6):615-620.
12. **Dey R, Bajpai V, Gandhi G, et al.** Application of artificial neural network technique for diagnosing diabetes melli-

tus. IEEE Third International Conference on Industrial and Information Systems, Kharagpur, India, 2008, 1-4.

13. **Heikes KE, Eddy DM, Arondekar B, Schlessinger L.** Diabetes risk calculator: a simple tool for detecting undiagnosed diabetes and pre-diabetes. *Diabetes Care* 2008. 31:1040-1045.

14. **Abbot RD.** Logistic regression in survival analysis. *Am J Epidemiol* 1985. 121(3):465-471.

15. **Shanker MS.** Using neural networks to predict the onset of diabetes mellitus. *J Chem Inf Comput Sci* 1996. 36:35-41.

16. **Kordos M, Blachnik M, Strzempa D.** Do we need whatever more than k-NN? In: Proceedings of the 10th International Conference on Artificial Intelligence and Soft Computing, Part I, Springer-Verlag Berlin, 2010, 414-421.

17. **Ster B, Dobnikar A.** Neural networks in medical diagnosis: comparison with other methods. In: Proceedings of the International Conference on Engineering Applications with Neural Networks, London, 1996, 427-430.

18. **Patil BM, Joshi RC, Toshniwal D.** Hybrid prediction model for type-2 diabetic patients. *Exp Syst Appl* 2010. 37:8102-8108.

19. **Jantan H, Hamdan AR, Othman ZA.** Human talent prediction in HRM using C4.5 classification algorithm. *Int J Comp Sci Engin* 2010. 2:2526-2534.

20. **Kohonen T.** Self-organizing formation of topologically correct feature maps. *Biol Cybern* 1982. 43:59-69.

21. **Fritzke B.** A growing neural gas network learns topologies. *Adv Neural Inf Process Syst* 1995. 7:625-632.

22. **Fritzke B.** Growing cell structures - a self-organizing network for unsupervised and supervised learning. *Neural Netw* 1994. 7:1441-1460.

23. **Deng D, Kasabov N.** On-line pattern analysis by evolving self-organizing maps. Proceedings of the 5th Biannual Conference on Aritificial Neural Networks and Expert Systems (ANNES), Dunedin, 2001, 46-51.

24. **Ali S, Abraham A.** An empirical comparison of kernel selection for support vector machines. 2nd International Conference on Hybrid Intelligent Systems, Soft Computing systems: Design, Management and Applications, IOS Press, The Netherlands, 2002, 321-330.

25. **Barakat NH, Bradley AP, Barakat MB.** Intelligible support vector machines for diagnosis of diabetes mellitus. *Trans Inf Technol Biomed* 2010. 14:1114-1120.

26. **Gunes PK, Aslan A.** A cascade learning system for classification of diabetes disease: generalized discriminant analysis and least square support vector machine. *Exp Syst Appl* 2008. 34:214-221.

27. **Purnami SW, Embong A, Zain JM.** A New smooth support vector machine and its applications in diabetes disease diagnosis. *J Comp Sci* 2009. 5:1006-1011.

28. **Lee YJ, Mangasarian OL.** A smooth support vector machine. *J Comp Optim Appl* 2001. 20:5-22.

29. **Basheer IA, Hajmeer M.** Artificial neural networks: fundamentals, computing, design, and application. *J Microbiol Meth* 2000. 43:3-31.

30. **Brent RP.** Fast training algorithms for multi-layer neural nets. *IEEE Trans Neural Netw* 1991. 2(3):346-354.

31. **Gori M, Tesi A.** On the problem of local minima in backpropagation. *IEEE Trans Pattern Anal Mach Intell* 1992. 14:76-85.

32. **Jaafar, SF, Ali DM.** Diabetes mellitus forecast using artificial neural networks. Asian conference of paramedical research proceedings, Kuala Lumpur, Malaysia, 2005, 5-7.

33. **Eskaf EK, Badawi O, Ritchings T.** Predicting blood glucose levels in diabetes using feature extraction and artificial neural networks. Third ICTTA conference, Damascus, 2008, 1-6.

34. **Jayalakshmi T, Santhakumaran A.** A novel classification method for classification of diabetes mellitus using artificial neural networks. International Conference on Data Storage and Data Engineering, Bangalore, 2010, 159-163.

35. **Chen LH, Chang S.** An adaptive learning algorithm for principal component analysis. *IEEE Trans Neural Netw* 1995. 6:1255-1263.

36. **Hagan MT, Menhaj M.** Training feed forward networks with the Marquardt algorithm. *IEEE Trans Neural Netw* 1994. 5:989-993.

37. **Gulbag A, Temurtas F.** A study on quantitative classification of binary gas mixture using neural networks and adaptive neuro fuzzy inference systems. *Sens Actuators B Chem* 2006. 115:252-262.

38. **Kayaer K, Yildirim T.** Medical diagnosis on Pima Indian diabetes using general regression neural networks. Proceedings of the international conference on artificial neural networks and neural information processing, Istanbul, 2003, 181-184.

39. **Temurtas H, Yumusak N, Temurtas F.** A comparative study on diabetes disease diagnosis using neural networks. *Expert Syst Appl* 2009. 36:8610-8615.

40. **Specht DF.** Probabilistic neural networks. *Neural Netw* 1990. 3:109-118.

41. **Buhmann, Martin D.** Radial basis functions: theory and implementations. Cambridge University Press, 2003. pp 54-78.

42. **Hagan MT, Demuth HB, Beale M.** Neural network design. PWS Publishing Company, 1996. pp 102-108.

43. **Bart K.** Neural networks and fuzzy systems: a dynamical systems approach to machine intelligence. Prentice Hall, 1992. pp 36-49.

44. **Polat K, Gunes S, Arslan A.** A cascade learning system for classification of diabetes disease: Generalized discriminant analysis and least square support vector machine. *Exp Syst Appl* 2008. 34:482-487.

45. **Dogantekin E, Dogantekin A, Avci D, et al.** An intelligent diagnosis system for diabetes on Linear Discriminant Analysis and Adaptive Network Based Fuzzy Inference System: LDA-ANFIS. *Digit Signal Process* 2009. 20:1248-1255.

46. **Smith JW, Everhart JE, Dickson WC, et al.** Using the ADAP learning algorithm to forecast the onset of diabetes mellitus. Proceedings of the Symposium on Computer Applications and Medical Care, IEEE Computer Society Press, 1988, 261-265.

47. **Carpenter GA, Markuzon N.** ARTMAP-IC and medical diagnosis: instance counting and inconsistent cases. *Neural Netw* 1998. 11:323-336.

48. **Jacobs RA, Jordan MI, Nowlan SJ, et al.** Adaptive mixtures of local experts. *Neural Comput* 1991. 3:79-87.

49. **Ubeyli ED.** Modified mixture of experts for diabetes diagnosis. *J Med Syst* 2009. 33:299-305.

50. **Chen K.** A connectionist method for pattern classification with diverse features. *Pattern Recognit Lett* 1998. 19:7545-558.

51. **Kamruzzaman SM, Hasan AR, Siddiquee AB, et al.** Medical diagnosis using neural network. Proceedings of the 3rd International Conference on Electrical and Computer Engineering, Dhaka, Bangladesh, 2004. 537-540.

262      **The Review of** Diabetic Studies             Shankaracharya, Odedra, et al.

Vol. 7 · No. 4 · 2010

52. **Ubeyli ED.** Automatic diagnosis of diabetes using adaptive neuro-fuzzy inference systems. *Expert Syst* 2010. 27:259-266.

53. **Stoean C, Stoean R, Preuss M, et al.** Diabetes diagnosis through the means of a multimodal evolutionary algorithm. Proceedings of the 1st East European Conference on Health Care Modelling and Computation, Craiova, Romania, Craiova, Medical University Press, 2005, 277-289.

54. **Shang N, Breiman L.** Distribution based trees are more accurate. Proceedings of ICONIP 96, Springer, Hong Kong, 1996, 133-138.

55. **Polat K, Gunes S.** An expert system approach based on principal component analysis and adaptive neuro-fuzzy inference system to diagnosis of diabetes disease. Digit Sign Proc 2007. 17:702-710.

56. **Friedman N, Geiger D, Goldszmit M.** Bayesian networks classifiers. *Mach Learn* 1997. 29:131-163.