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Automatic Classification of Colorectal and Prostatic Histologic tumor Images using Multiscale Multispectral Local Binary Pattern Texture Features and Stacked Generalization

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Abstract

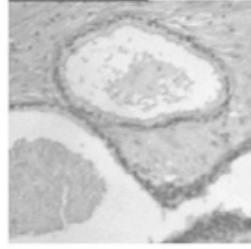
This paper proposes a new multispectral multiscale local binary pattern feature extraction method for automatic classification of colorectal and prostatic tumor biopsies samples. A multilevel stacked generalization classification technique is also proposed and the key idea of the paper considers a grade diagnostic problem rather than a simple malignant versus tumorous tissue problem using the concept of multispectral imagery in both the visible and near infrared spectra. To validate the proposed algorithm performances, a comparative study against related works using multispectral imagery is conducted including an evaluation on three different multiclass datasets of multispectral histology images: two representing images of colorectal biopsies - one dataset was acquired in the visible spectrum while the second captures near-infrared spectra. The proposed algorithm achieves an accuracy of 99.6% on the different datasets. The results obtained demonstrate the advantages of infrared wavelengths to capture more efficiently the most discriminative information. The results obtained show that our proposed algorithm outperforms other similar methods.

Keywords: Multiscale Multispectral Local Binary Pattern, Stacked generalization, histology, colorectal cancer, prostate cancer, automatic diagnosis

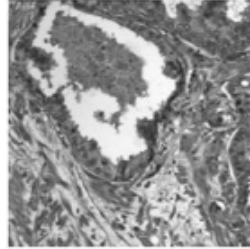
1. Introduction

The World Health Organization has declared that the cancer burden is a worldwide health problem. According to their 2014's report, 14 million new cases were diagnosed in 2012 and 8 million people died from it in the same period [1]. Colorectal cancer is the third most incident globally and prostate is in second position amongst men representing respectively 9.7% and 7.9% of all cancers for both sexes [1]. Both colorectal and prostatic tissues are glandular thus having a similar histological appearance. They also are both subject to the same tumor types; adenocarcinoma being the most commonly diagnosed cancerous tumor type in these organs. The European Association of Urology's guidelines [2] advise to perform a biopsy and a histological analysis on the sample for prostate cancer diagnosis. This method is also the most widely used for colorectal cancer diagnosis [3]. However, this process is very time-consuming for pathologists as they have to manually analyze every sample to spot the particular features characterizing the type of tumor and the various cancer stages. This process results in a high intra- and inter-observer variability [4], [5] thus affecting the diagnostic reliability. This paper aims to propose an algorithm that will automatically classify the samples into different categories of the cancer hence assisting the pathologist to make the appropriate diagnostics. This will in turn help to reduce the diagnosis time and act as a second opinion for the pathologist to reduce the intra- and inter-observer diagnostic variability.

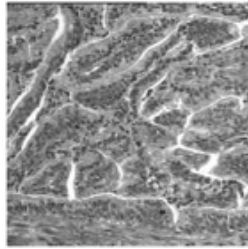
Fig. 1 shows microscopic images of three different colorectal tumor biopsies. The first one is a Benign Hyperplasia, a benign tumor with little risk of evolving to cancer while the second represents an Intraepithelial Neoplasia, a tumor with high risk of evolving to cancer. The last image shows a carcinoma, which is a cancerous tumor. One of the features taken into account by the pathologists for the diagnosis is the general structure and organization of the tissue. In a normal structure, the epithelial cells are organized around the lumen which is separate from the plasma cells; whereas in the case of a carcinoma the normal structure becomes completely random and can be chaotic. Therefore, the proposed algorithm uses image texture features so as to capture and quantify its structure in order to classify the samples into the various types of malignancy. The sole pixel intensity can be insufficient to characterize the type of cell or sub-cellular components and so can have



(a) Benign Hyperplasia



(b) Intraepithelial Neoplasia



(c) Carcinoma

Figure 1: Example of images from the different classes of colorectal tissue.

negative effect on the feature extraction. Consequently, using the spectral response of each point of the sample to describe the tissue is adopted in this paper to improve the classification performances; hence the use of multispectral images of the biopsies. This paper also aims to investigate the advantages of using the pixels response from a wider electromagnetic spectrum ranging from the visible light to the infrared (IR) in comparison to other methods that can be used in biopsy image analysis as shown in Fig. 2.

The main contributions of this paper are three-fold. First, our work considers a grade diagnostic problem rather than a simple malignant versus benign tumor problem in the context of multispectral imaging and this is almost absent in the literature, especially for colorectal cancer [6]. Second, it introduces a new Multispectral Multiscale Local Binary Pattern (MMLBP) texture feature which is an adjusted LBP to multispectral data taking into account the third dimension (spectrum) of the data. The MMLBP differs from the traditional LBP in that it considers the joint information across spatial and spectral directions of the image. In addition, a stacked generalization technique is devised in order to fuse the different scales of the MMLBP and GLCM features at the score level. Finally, a new dataset is also introduced in

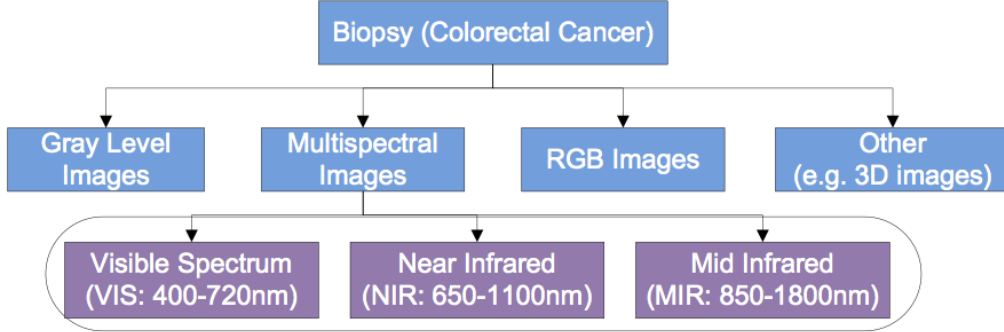


Figure 2: Imaging technologies for the classification of colorectal cancer.

the paper. This dataset is composed of multispectral images with a spectrum extending to the infrared (IR).

This paper is organized as follows: Section 2 gives a briefly a review of existing systems including a discussion on prostate and colorectal cancer tissue analysis . Section 3 reports some related feature extractors using LBP approach and some of its variants. Section 4 describes the proposed methodology including the contributions made and the proposed system. Section 5 explains the implementations and experimental analysis. Section 6 evaluates the results and a comparative study against existing techniques. It also shows the advantages of using IR images to improve the classification accuracy. Section 7 concludes the paper.

2. Related work

2.1. Previous work on prostate and colorectal cancer tissue analysis

Several techniques available in the literature extract textural features using panchromatic images [6], [7]. Esgiar *et al.* [8] computed a gray-level co-occurrence matrix (GLCM) on each colorectal histological image and extracted some of the GLCM features [9]. They then proposed a malignant versus benign classification using an SVM classifier. In [10], Kalkan at al. combined the same features with structural ones before computing a feature selection and a four-class classification, and achieved a 75.15 % accuracy. In [11], the authors used a 8-class dataset of 5000 images. The classes involved where the following: tumor epithelium, simple stroma, complex stroma, immune cells, debris, normal mucosal glands adipose tissue and background (no

tissue). The authors compared several texture descriptors such as GLCM, LBP, perception feature - mimicking the human perception at an abstract level - and Gabor filters. Their results range from 74 to 97 %. In the case of prostate cancer, several authors used the GLCM features for the same task of carcinoma detection [12]. The authors of [13], [14] used fractal analysis for prostate cancer grading or carcinoma detection.

Multispectral images have been used for texture feature extraction. In [15], Masood *et al.* applied GLCM features after segmenting the image data through a pre-processing phase. the approach consists of using the spectral dimensions to segment the image into four clusters representing four different tissue types: nuclei, cytoplasm, glands and stroma. Chaddad *et al.* proposed an improved version of the snake algorithm for the segmentation and extraction of GLCM texture features of multispectral-segmented images [16]. In [17], the authors proposed a method for characterizing the continuum of colorectal cancer using several texture features after segmentation. As features extraction, the GLCM features, the Laplacian of Gaussian and discrete wavelets were used. A few other studies used wavelet transforms [18] [19] and Laplacian of Gaussian [20] [10]. In [21], Roula *et al.* worked on prostate histological images and extracted GLCM features from each spectral band and combined them with morphological features for the discrimination phase using a quadratic discriminant analysis. They showed that multispectral analysis significantly improved the classification scores. The authors of [22] also demonstrated that the use of texture features in multispectral images improved the results when using texture features of panchromatic images on a colorectal histology dataset. They compared the performance of different texture features on multispectral images, namely the GLCM features and the multiscale LBP and used PCA for dimensional reduction followed by a SVM classifier. In [23], Tahir *et al.* first extracted statistical and structural features as well as the GLCM features. They then used a Round-Robin Tabu Search for dimensional reduction of the multispectral data before classification. They achieved a classification accuracy between 98% and 100%.

None of the previously mentioned authors used a multispectral texture feature detector that uses the spectral dimension directly. They either combined several results of 2-dimensional texture detector run on each spectral band, or used the dimensional reduction to create a 2D image on which the texture was to be detected. Khelifi *et al.* [24] developed a multi-band texture detection extending the GLCM. For this purpose, they used a spatial and spectral gray level dependence method (SSGLDM) assuming a joint infor-

115 mation between spectral bands exists. They applied this technique to the
116 prostate cancer case.

117 However, only few studies use LBP texture features in this field [22, 25, 26]
118 and none of them uses the joint information of spatial and spectral dimen-
119 sions. For example, the authors of [25] select a single band from which the
120 texture extraction was conducted. In [26], the LBP histogram is built on all
121 three color channels of the image.

122 2.2. Previous work on multispectral texture analysis

123 Some methods for other applications, such as image segmentation, used
124 a 3D histogram as a mean to fuse information from three color channels
125 of a colour image [27]. Hassan El Maia *et al.* [28] proposed a method for
126 multispectral image classification using the mutual information of GLCM
127 features. In [29], the authors used a method developed in [30] for automatic
128 face recognition. This algorithm was a modified LBP that computed a LBP
129 on each color band of the spectrum separately and added opponent features
130 to capture the spacial correlation between the bands. Radu-Mihai Coliban
131 *et al.* [31] proposed a pseudo-morphology based on the Euclidean distance
132 in \mathbb{R}^n . Using the proposed pseudo-morphology, the authors introduced a
133 pseudo-granulometry and a morphological covariance to characterize the im-
134 age texture. In [32], the authors use a neural network structure to classify
135 multispectral texture information extracted from the images.

136 2.3. Previous work on IR texture analysis

137 In the field of facial recognition, the IR spectrum has been used and
138 has proven to increase the recognition rates in many cases. Abdelhakim
139 Bendada *et al.* [33] introduced a differential LTP descriptor and extend their
140 method to the IR spectrum. They showed that a high recognition rate was
141 achieved with the IR spectrum. The authors [34] developed a method for
142 synthesising the visible and near IR face images in order to take advantage
143 of both the illumination invariance of IR images and the detailed texture
144 information provided by the face images captured in the visible range of
145 the electromagnetic spectrum. The authors compared their method to the
146 conventional LBP applied separately to the near IR and the visible images
147 and showed that the combined use of the IR and the visible spectra increased
148 the identification rate by 8.76pp (from 88.83% to 97.59%).

149 Thematic mapping imagery uses the infrared spectrum to acquire infor-
 150 mation not captured by the visible spectrum. Yun Zang [35] used an algo-
 151 rithm of conditional variance detection on multispectral images captured on
 152 a visible and IR spectrum for classification of urban treed areas.

153 3. Feature extraction using LBP approach: a review

154 In this section, the conventional LBP and its rotation invariant and three-
 155 dimensional variants are discussed.

156 3.1. Conventional LBP

157 Ojala *et al.* described LBP texture features as a local characterisation
 158 of a pixel's neighborhood at a radius R sampled into a set of P neighbors
 159 on a circle centered around the central pixel and of radius R . Let g_0 be the
 160 intensity of the central pixel x and g_p the intensity of its p^{th} neighbor. The
 161 LBP is defined as follows [36]:

$$LBP_{P,R}(x) = \sum_{p=1}^P s(g_0 - g_p) 2^{p-1} \quad (1)$$

where,

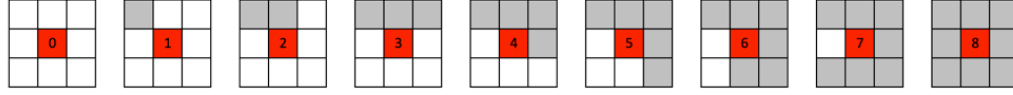
$$s(x) = \begin{cases} 0 & \text{if } x \leq 0 \\ 1 & \text{if } x > 0 \end{cases}$$

162 LBP is computed for the whole image, before it is pooled into a LBP his-
 163 togram of size 256. The resulting LBP histogram, which is invariant to
 164 intensity changes, is then used as a texture feature descriptor to characterize
 165 the image.

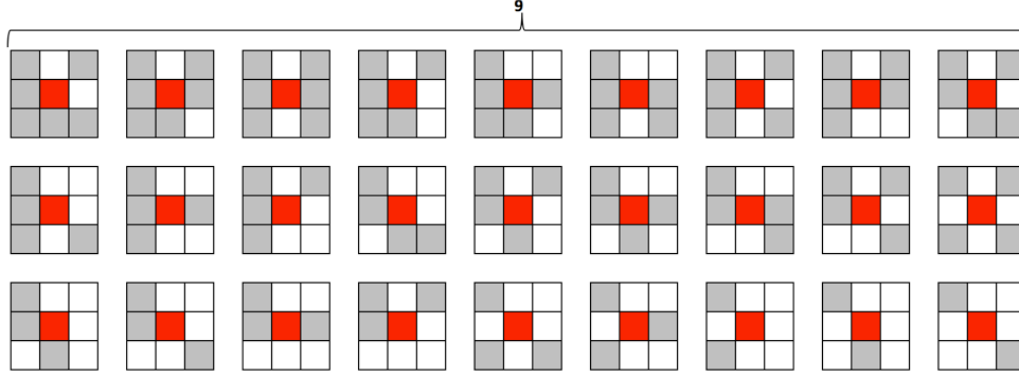
166 3.2. Rotation Invariant Uniform LBP

167 A rotation invariant LBP, referred to as LBP^{riu2} , using uniform patterns
 168 has also been proposed as illustrated in Fig. 3a. They operate as templates
 169 for microstructures such as bright spot (0), flat area or dark spot (8) and
 170 edges of varying positive or negative curvature (1-7) [36]. These structures
 171 define a uniformity measure U corresponding to the number of transition in
 172 the pattern as follows:

$$U(LBP_{P,R}) = |s(g_{P-1} - g_c) - s(g_0 - g_c)| + \sum_{p=1}^{P-1} |s(g_p - g_c) - s(g_{p-1} - g_c)| \quad (2)$$



(a) Uniform LBP patterns and their corresponding labels



(b) Non-uniform LBP patterns

Figure 3: The 36 unique possibilities for a circular symmetric set of LBP patterns and their corresponding labels for rotation invariant, uniform LBP. The red squares correspond to the central pixel, the white and grey squares represent the 0 and 1 bits in the 8-bits output of the operator. The numbers are the unique $LBP_{P,R}(x)$ labels.

Fig. 3a shows the 9 patterns with a U measure of at most 2 when the 27 other patterns shown of Fig. 3b have a uniformity measure of at least 4. Therefore, patterns having $U(LBP_{P,R}) \leq 2$ are said to be uniform. The following operator defines a gray-scale and rotation invariant texture description [36]:

$$LBP_{P,R}^{riu2}(x) = \begin{cases} \sum_{p=1}^P s(g_p - g_c) & \text{if } x \leq 2 \\ P + 1 & \text{otherwise} \end{cases} \quad (3)$$

In this way, $P + 1$ uniform patterns are assigned to a unique label corresponding to the number of 1 bits in the pattern while the non-uniform patterns are grouped under the same category. The final texture feature used is a histogram of $P + 2$ bins generating all the $LBP_{P,R}^{riu2}$ outputs accumulated over the image.

This form of LBP seems more adapted to the problem at hand because of the rotation invariance it provides. Indeed, in the case of histopathology, sample orientation and cells direction are not relevant criteria to consider for classification because they vary independently to the sample's class. A

187 second advantage of this $LBP_{P,R}^{riu2}$ over a conventional $LBP_{P,R}$ is its smaller
 188 size thus making it faster to process in a classification step.

189 3.3. Three-dimensional LBP

190 Since multispectral images are three-dimensional data the conventional
 191 LBP concept needs to be modified to deal with this datatype. In the lit-
 192 erature, two methods are usually described when dealing with 3D images
 193 for applications such as video processing and face recognition [37]. The
 194 proposed method is inspired from Volume LBP and LBP-TOP (for Local
 195 Binary Pattern-Three Orthogonal Plan) [37]. Here, we briefly discuss VLBP
 196 and LBP-TOP before explaining the proposed extension of LBP to multi-
 197 spectral LBP. To extend LBP to Dynamic Texture analysis, Zhao *et al.* define
 198 a neighborhood as the joint distribution of $3P + 3$ image pixels where P is
 199 the number of neighbors on one frame as shown on [37]. A similar technique
 200 to the conventional LBP can be applied and a Volume Local Binary Pattern
 201 (VLBP) is defined as follows:

$$VLBP_{P,R}(x) = \sum_{p=1}^{3P+2} s(g_0 - g_p)2^{p-1} \quad (4)$$

202 The VLBP local features are pooled into a histogram of size 2^{3P+2} . This
 203 histogram's size increases very quickly when the number of neighbors P grows
 204 and may become very computationally intensive. On the other hand, using
 205 a small P may lead to a loss of some critical information for diagnosis. To
 206 address this issue, a LBP-TOP feature is proposed by considering three or-
 207 thogonal planes intersecting on a central pixel as shown in [37]. The technique
 208 computes a two-dimensional LBP on each of these plans and concatenates
 209 the output histograms which will be of size $3 * 2^P$ instead of 2^{3P+2} in the
 210 previous case. Here, the circles are considered in the time dimension because
 211 this LBP-TOP is meant to be applied on video processing so the motion
 212 direction of texture is unknown.

213 4. The Proposed MMLBP System for Cancer Classification

214 As illustrated in Fig. 4, the proposed system is composed of two main
 215 stages. First, MMLBP features are extracted and, then, an Independent
 216 Component Analysis is performed to reduce the dimensionality of the feature
 217 space. Then, a stacked generalization employing the Support Vector Machine
 218 classifier is used at the matching stage.

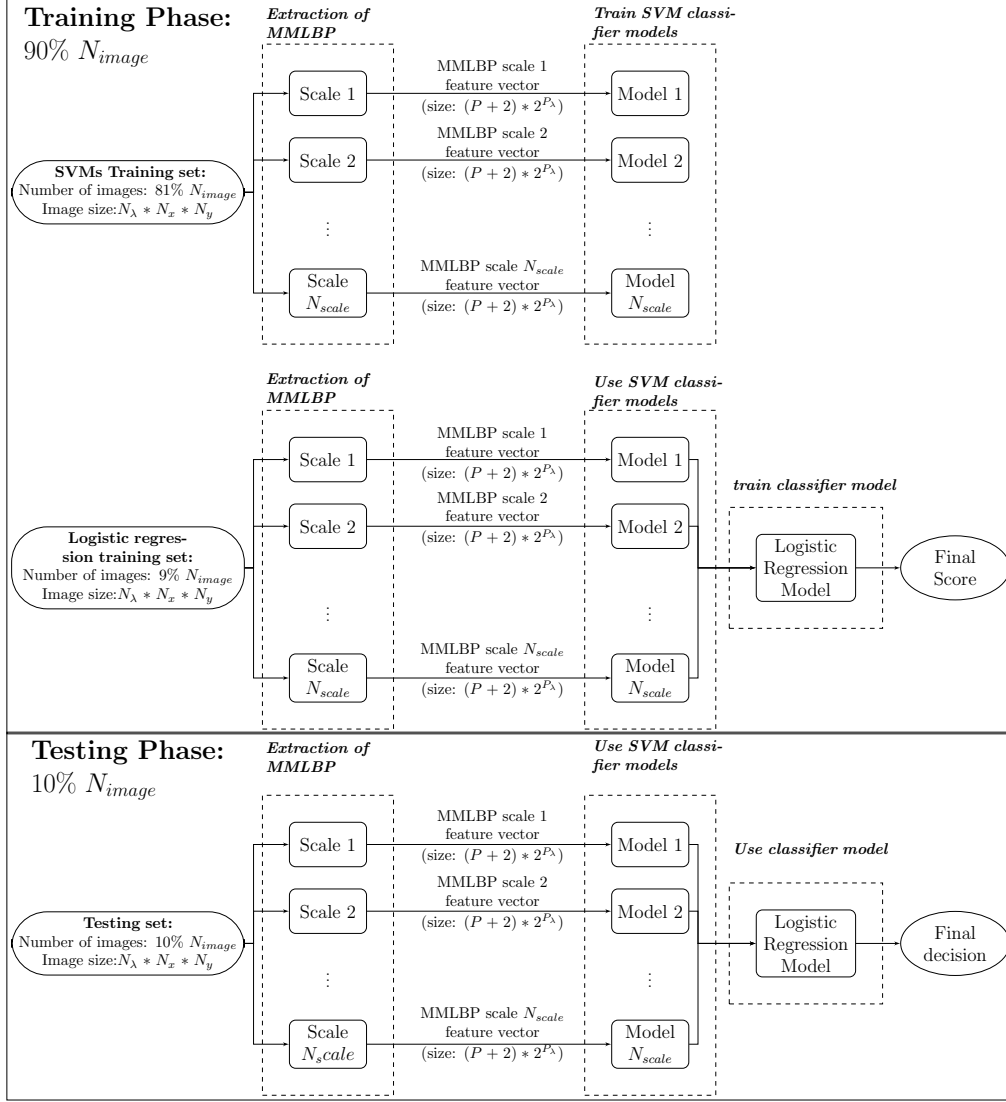


Figure 4: Block diagram of stacking training and testing with MMLBP texture features. N_{image} represents the number of images in the dataset; N_x , N_y are the number of lines and columns in each image, respectively, and N_λ is the number of spectral bands.

4.1. Proposed Multispectral LBP Texture Feature

In the proposed technique the third dimension is spectral (not temporal) thus no texture motion is considered. Therefore, unlike in the aforementioned 3D-LBP variants, a neighborhood of only P points in the spatial plan and P_λ

223 on a straight line in the spectral dimension intersecting the spatial plan at the
 224 central pixel was considered as shown in Fig. 5 with $P_\lambda = 2$. As explained
 225 above, this technique is adopted to make the LBP rotation invariant in the
 226 spatial dimensions while still using the same U measure described in (3)
 in the XY plan. The idea is to assign the $LBP_{P,R}^{riu2}$ patterns to different

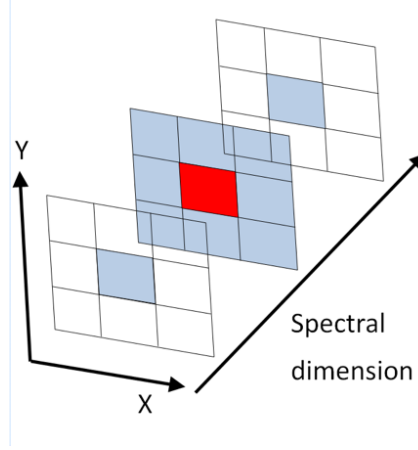


Figure 5: Multispectral LBP descriptor: the neighborhood considered for multispectral LBP. X and Y being the spatial dimensions. Each tile represents a pixel. The red tile is the central pixel considered, and the blue tiles are the pixel considered in the neighborhood.

227 categories depending on the P_λ pixels in the neighboring plans. On top of
 228 the $LBP_{P,R}^{riu2}$ computed using equation 3, $LBP_{P_\lambda,R}^\lambda$ is calculated using the
 229 following equation:
 230

$$LBP_{P_\lambda,R}^\lambda(x) = \sum_{q=1}^{P_\lambda-1} s(g'_q - g_c)2^q \quad (5)$$

231 where, g'_q is the pixel value in the pixel of plan q aligned to the central pixel.
 232 The $MMLBP_{P,P_\lambda,R}$ is defined as follows:

$$MMLBP_{P,P_\lambda,R} = LBP_{P,R}^{riu2} + (P+1)LBP_{P_\lambda,R}^\lambda \quad (6)$$

233 The $MMLBP_{P,P_\lambda,R}$ outputs are then pooled into a histogram of size
 234 $(P+2) * 2^{P_\lambda}$. It is worth noting that the scale is controlled by $R \in [1..N_{scale}]$
 235 As a result, the histograms built from each scale are concatenated to form
 236 the MMLBP and each scale is considered as a separate feature which is fed
 237 to the stacked classifier as shown in Fig. 6.

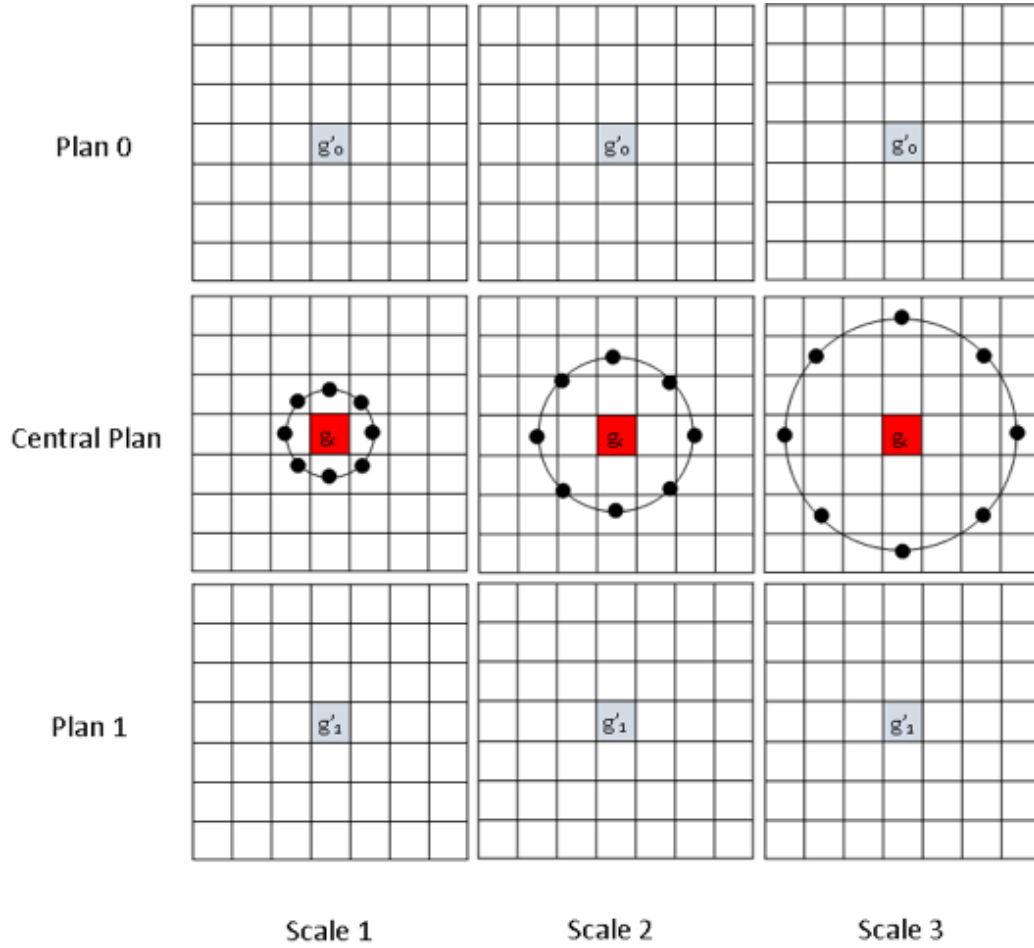


Figure 6: Multiscale neighborhood for MMLBP

238 *4.2. Dimensionality Reduction using Independent Component Analysis and*
239 *Classification using Support Vector Machine*

240 In order to address the curse of dimensionality problem and hence reduce
241 the learning cost, the Independent Component Analysis (ICA) is applied
242 before classification. In contrast to more widely used Principal Component
243 Analysis (PCA), this technique presents the advantage of being able to de-
244 correlate the signal and reduce statistical dependencies between the features
245 as much as possible [38]. In fact, it could be seen as a version of PCA that
246 defines orthogonal directions. The ICA transformed data are computed using
247 only the training data of the SVM classifier. The testing data are projected

248 to the new basis before classification. The number of components used for
 249 classification will be optimized as described in a further section.

250 The classification step consists of a multiclass Support Vector Machine
 251 (SVM) classifier with a Gaussian kernel. A SVM constructs a hyper-plan
 252 separating the classes; it tries to find the maximum distance to the nearest
 253 training data point of each class and can be described as follows. Given a
 254 training vector $x_i \in \mathbf{R}^P$, $I \in [1, n]$, in two classes and a vector $y \in \{1, -1\}^n$,
 255 the SVM solves the following optimization problem:

$$\min_{w,b,\zeta} (1/2 w^T w + C \sum_{i=1}^n \zeta_i) \text{ subject to: } y_i(w^T \phi(x_i) + b) \geq 1 - \zeta_i, \zeta_i \geq 0, i \in [1, n] \quad (7)$$

256 Its dual form is:

$$\min_{\alpha} (1/2 \alpha^T Q \alpha - e^T \alpha) \text{ subject to: } y^T \alpha = 0, 0 \leq \alpha_i \leq C, i \in [1, n] \quad (8)$$

257 Where, $C > 0$ is the upper bound, Q is an n by n positive semi definite
 258 matrix $Q_{i,j} = y_i y_j K(x_i, x_j)$ where $K(x_i, x_j) = \phi(x_i)^T \phi(x_j)$ is the kernel. Here
 259 training vectors are implicitly mapped into a higher dimensional space by the
 260 function ϕ . The decision function is:

$$f_{decision} = \text{sgn}(\sum_{i=1}^n y_i \alpha_i K(x_i, x) + \rho) \quad (9)$$

261 The kernel function used here is the radial basis function or Gaussian
 262 kernel:

$$K(x) = e^{-\gamma |x - x'|^2} \quad (10)$$

263 Where γ is a positive parameter. The kernel parameters are optimized using
 264 a grid search method which will be detailed in Section 5.1. In order to
 265 find the appropriate compromise between the sizes of training and testing
 266 datasets and hence avoid over-fitting that might be caused by a leave-one-
 267 out technique; a 10-fold cross-validation is used. The one-versus-all technique
 268 is used to build the multiclass classifier.

269 4.3. Logistic Regression for Stacked Generalization

270 Stacked generalization (or stacking) is an ensemble method for classifica-
 271 tion [39]. It uses the output of a first layer of classifiers as inputs to another

272 classifier - called meta-classifier - for the final decision. In this paper, this sys-
 273 tem is used to fuse the different scales of multispectral LBP texture feature
 274 at score level.

275 Fig. 4 shows the two steps of training and testing for the stacking algo-
 276 rithm. A logistic regression model is used as a meta-classifier for its relatively
 277 low computing cost. The first layer of classifiers is composed by SVM clas-
 278 sifiers with a Gaussian kernel as described in Section 4.2. In addition to a
 279 10-fold cross-validation carried out at the meta-classifier level, an internal
 280 cross-validation of the training data is implemented in order to prevent bias
 281 and improve stability of the different classifiers.

282 5. Experiment and Setup

283 5.1. Datasets

284 To evaluate the performance of the proposed technique, three different
 285 datasets are used in the experimentation process.

286 5.1.1. Dataset 1: Colorectal tumor Tissue from Texas

287 The first one, described in [17], is composed of colorectal biopsy images
 288 acquired using multispectral imagery at low magnification power (x40). The
 289 database consists of 29 three-dimensional images having a spatial resolution
 290 of 512*512 pixels and 16 spectral bands corresponding to wavelengths be-
 291 tween 500 and 650 nm. The images are divided into 3 classes of tumor tissue
 292 types: Carcinoma (Ca), the class containing the cancerous samples, Benign
 293 Hyperplasia (BH), a class with benign tumors, and Intraepithelial Neoplasia
 294 (IN), containing images with tissues at a precancerous stage.

295 5.1.2. Dataset 2: Prostatic tumor Tissue

296 The second dataset [23], with some samples shown in Fig. 7, consists
 297 of multispectral images taken at 16 spectral channels (from 500 to 650 nm)
 298 and at x40 magnification power. 592 different samples (multispectral images)
 299 of size 128*128 have been used to carry out the analysis. The samples are
 300 evaluated by two highly experienced independent pathologists and labeled
 301 into four classes: 165 cases of Stroma (Str), which is normal muscular tissue,
 302 106 cases of Benign Prostatic Hyperplasia (BPH), a benign condition, 144
 303 cases of Prostatic Intraepithelial Neoplasia (PIN), a pre-cancerous stage, and
 304 177 cases of Prostatic Carcinoma (PCa), an abnormal tissue development
 305 corresponding to cancer.

5.1.3. Dataset 3: Colorectal Vis-IR

Colorectal Vis-IR: The third dataset is also composed of multispectral colorectal histology data with a (x40) magnification power. This dataset was developed by University of Qatar with the collaboration of the Al-Ahli Hospital, Doha and will be made available for public use¹. It is split into 4 classes, each of them composed of 10 images. The images are acquired on a wider spectrum than in the first dataset as it is spread on the visible (Vis) and infrared (IR) ranges of the electromagnetic spectrum - shown in Figure 2 - with an interval of 23 nm between each wavelength. That is to say, in the visible range, the wavelength interval is 23 nm starting from 465 nm to 695 nm and in the IR range, the wavelength interval is also 23 nm and ranges from 900 nm to 1590 nm. The 4 classes are: Carcinoma (Ca), containing the images of cancerous colon biopsies, Tubular Adenoma (TA), a pre-cancerous stage, Hyperplastic Polyp (HP), a benign polyp and No Remarkable Pathology (NRP).

5.2. Experiments

The first two datasets are used for the first sets of experiments and the 3rd dataset is used in the last experiment to show how the performance can be improved by using the IR imagery.

The proposed system is first compared with the results given by the algorithm described in [22] by using a conventional LBP extracted from a panchromatic image that is generated by averaging the spectral bands of the multispectral image. It is also compared to another variant of LBP adapted to multispectral images. It consisting in extracting LBP histograms from each band and then concatenating them to form a final descriptor. This method is referred to as the concatenated LBP. It is worth mentioning that these LBP variants were used with an SVM classifier for a fair comparison. For the same reason, they were also applied using the same number of scales N_{scale} . Many authors use GLCM texture features - see Appendix: GLCM texture feature. The results obtained using the proposed system - that is the stacked classification of the GLCM feature model combined with the MMLBP as shown in Figure 8, this is denoted as stacked MMLBP + GLCM - were also compared to the ones given by MMLBP alone. In order to assess our algorithm's robustness, the two first datasets presented in Section 5.1 are used.

¹it is expected that the first release will take place in January 2018

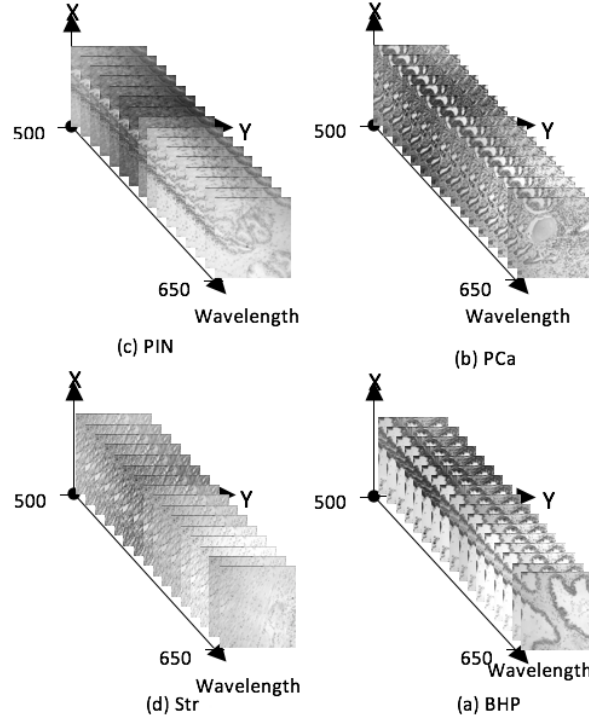


Figure 7: Sample of multispectral images from Dataset 2

340 In the second set of experiments, the impact of spatial resolution varia-
 341 tions in performance is addressed .

342 The algorithm is also compared against different algorithms from the
 343 literature. An adapted version of Masood *et al.*'s algorithm [15] to the mul-
 344 ticlass problem is implemented. In this method they use the GLCM features
 345 after segmentation of the image to train an SVM classifier. The results given
 346 by the algorithm described by the authors of [16] are used for comparison.
 347 It consists of using a snake algorithm for image segmentation and uses the
 348 GLCM feature as well. Our method is also compared against Khelifi *et al.*'s
 349 results [24]. They define a multispectral form of the GLCM before extracting
 350 the GLCM features. Finally the results shown in [23] are used for comparison
 351 purposes. In that paper, Tahir *et al.* describe a Round-Robin Tabu search
 352 algorithm for prostatic tumor classification.

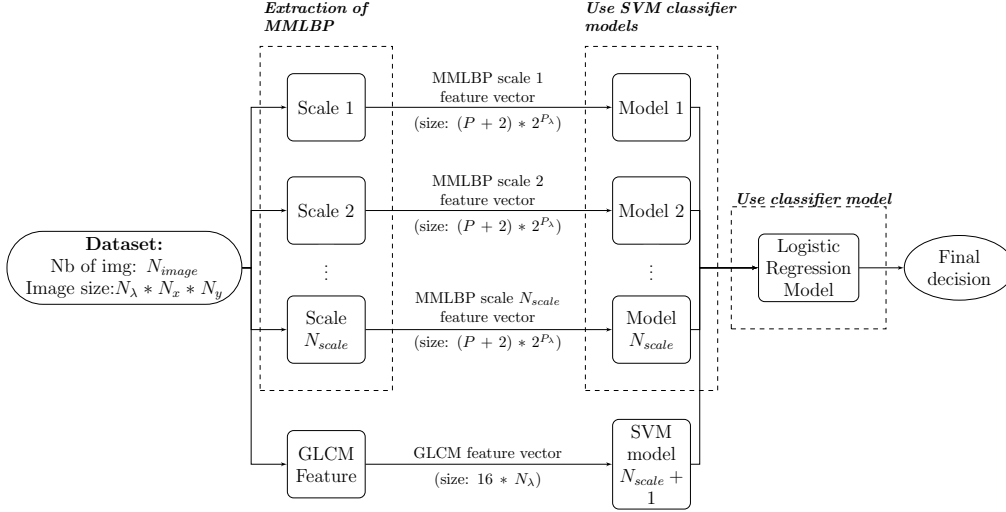


Figure 8: Block Diagram of the proposed algorithm.

5.3. Evaluation Measures

In order to avoid accuracy variations, the cross-validation is run ten times and the accuracy is averaged. The standard deviation is calculated on the mean accuracies of each cross-validation.

In addition of the accuracy and the standard deviation, the ROC curve and the Area Under Curve and the confusion matrix are also computed and used to assess the performances of the proposed algorithm. These performance measures are useful metrics to allow for a better understanding of what each class captures before one-versus-all combination to obtain the overall accuracy.

6. Results and Discussion

6.1. Training procedure

As illustrated by Fig. 4, the double 10-fold cross-validation run on the datasets means that, for each experiment, 90% of the dataset is used for training the logistic regression classifier model and the remaining 10% are used for the testing phase. 90% of this training set (or 81% of the total dataset) is used for training the SVM models and in the remaining 10% of the training set (or 9% of the whole dataset), the trained SVM models are

371 used to train the logistic regression model. Table 1 displays the SVMs and
 372 logistic regression training sets and the testing set sizes for each dataset.

Table 1: Number of images used in each phase for each the tested dataset

Data-set	SVMs training set	Logistic regression training set	testing set	dataset size
1	23	3	3	29
2	480	53	59	592
3	29	3	4	36

373 6.2. Parameters Tuning

374 As discussed previously, a total of 3 parameters need to be optimized for
 375 each SVM classification: the number of components selected in the ICA, and
 376 the C and γ parameters of the SVM kernel from Eq. (8), (10). A three-
 377 dimensional grid search was performed with the following parameters, with
 378 a step equals to 1:

$$\begin{aligned}
 C &= 10^i, \text{ with } i = [-3 : 3], \\
 \gamma &= 10^j, \text{ with } j = [-3 : 3], \\
 N_{comp} &= 10 * k, \text{ with } k = [1 : 50].
 \end{aligned}$$

379 For each combination of the parameters in these intervals, the accuracy is cal-
 380 culated and averaged with a 10-fold cross-validation. The parameters giving
 381 the maximum average accuracy are then chosen as the model parameters.

382 6.3. Proposed Algorithm Discussion

383 Table 2 shows a comparison of the classification accuracies obtained us-
 384 ing different features and classification methods. First a conventional LBP
 385 followed by a SVM classification is performed and an accuracy of 88.3 % is
 386 achieved on dataset 1 and 77.4 % on dataset 2. This shows this option is
 387 not robust to the data. When using a concatenated version of multispectral
 388 LBP followed by an SVM classification, the results are improved and an ac-
 389 curacy of 95.8 % is achieved on dataset 1. However, only 89 % accuracy is
 390 obtained on dataset 2 hence indicating the instability of the method. When
 391 using stacked generalization with MMLPB texture feature, the results are im-
 392 proved again and an accuracy of 99.0 % and 99.2 % on dataset 1 and dataset

2, respectively, thus demonstrating the robustness of the proposed algorithm. This can be explained because the stacking method selects the best features for classification and discards the features that drop the accuracy and this is independent to the data. When GLCM texture features are combined to the MMLBP texture features the results are improved by 0.3 - 0.6 percentage points (pp). It can also be seen that the multispectral information brings significant improvement over the conventional LBP as illustrated by the performance of the concatenated multispectral LBP method. Furthermore, the stacking classification process enhances the performance further as demonstrated by the results of the stacked LBP compared to the concatenated LBP.

Table 2: Accuracy comparison of different feature extraction and classification methods

Data-set	Conventional LBP (%)	Concatenated LBP (%)	Stacked MMLBP (%)	Proposed algorithm: Stacked MMLBP + GLCM (%)
1	88.3 ± 2.7	95.8 ± 0.5	99.0 ± 0.3	99.6 ± 0.4
2	77.4 ± 4.0	89.0 ± 0.9	99.2 ± 0.3	99.5 ± 0.3

Figure 9 and 10 displays the ROC curves and shows the Area Under Curve (AUC) for the different classes in a binary classification following the one versus all scheme. This is done to assess the positive and negative false alarm rates for each class. Table 3 and Table 4 show the confusion matrices obtained when using different datasets.

Table 3: Confusion Matrix for dataset 1

	Class BH	Class Ca	Class IN
Class BH	144	0	0
Class Ca	0	144	0
Class IN	1	4	139

As can be seen from Fig. 9, the system performs better on classes BH and Ca than it does on class IN. Fig. 6.3 displays some examples of correctly classified and misclassified images from dataset 1. Fig. 11a shows a sample of class IN which has been misclassified as class Ca by the proposed algorithm. Fig. 11b and 11c show correctly classified samples from class IN and Ca, respectively. As one can see the contrast on Fig. 11a is not as pronounced as

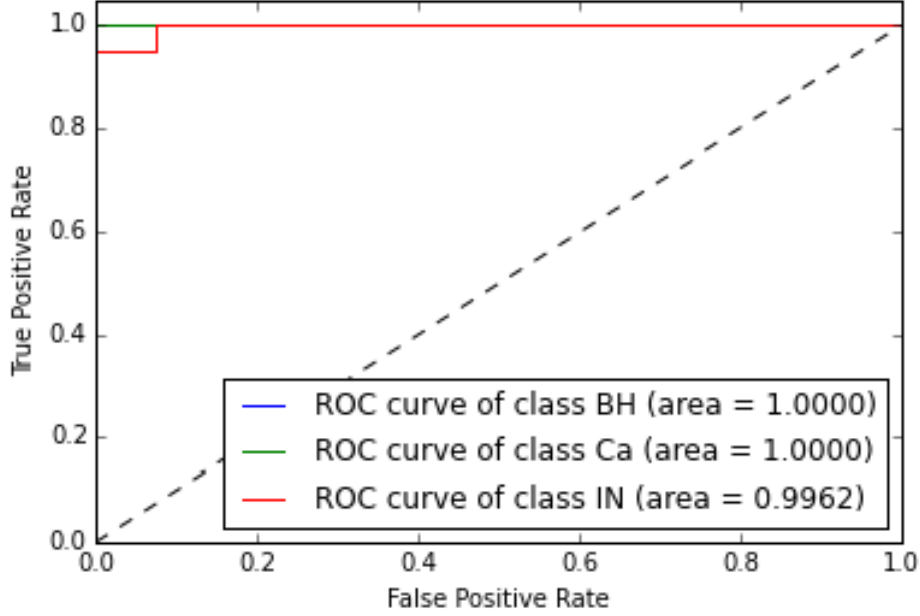


Figure 9: ROC for the proposed algorithm for Dataset 1

Table 4: Confusion Matrix for dataset 2

	Class BPH	Class PCa	Class PIN	Class Str
Class BPH	128	0	0	0
Class PCa	0	173	3	0
Class PIN	0	0	144	0
Class Str	0	0	0	144

415 what can be observed on Fig. 11b. This is especially true for the epithelial
 416 cells: in Fig. 11a the outer border of the cytoplasm of the cell is not as visible
 417 as it is on Fig. 11b. On the other hand, Fig. 11c presents hyperchromatism
 418 meaning the nuclei of the cells are well contrasted with the rest of the tissue
 419 and the border of the cytoplasm is not very clear. These features described
 420 on Fig. 11c is similar to what can be observed on Fig. 11a. Also both Fig.
 421 11a and 11c show an area with stroma tissue respectively at the bottom right
 422 and at the top left of the images. The combination of both these features
 423 can explain the misclassification of Fig. 11a.

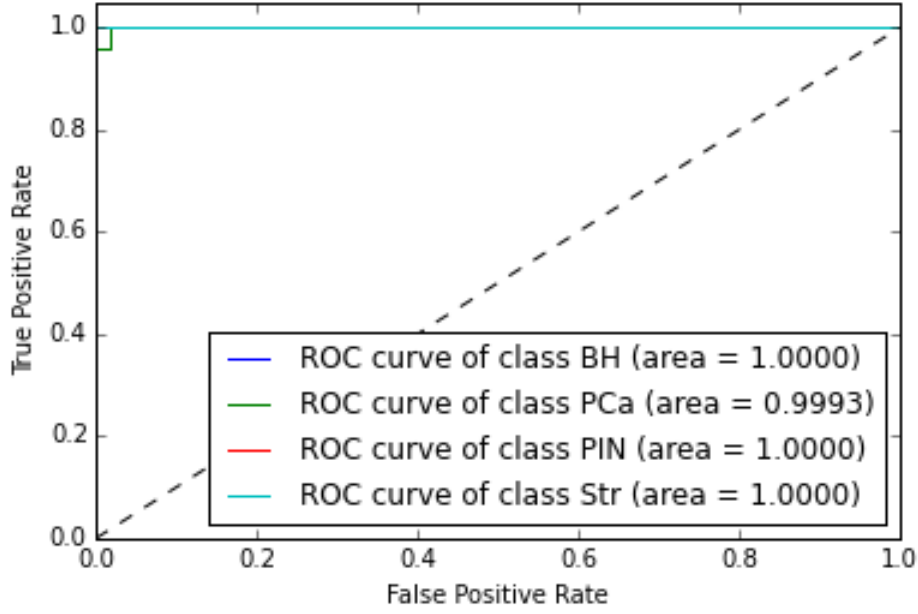


Figure 10: ROC for the proposed algorithm for Dataset 2

6.4. Impact of the Spatial resolution

Table 5 shows the impact of image spatial resolution on the results. As can be seen, the accuracy is marginally influenced by the change of resolution. It varies from $99.6\% \pm 0.4$ for the full resolution to $98.7\% \pm 0.4$ for a spatial resolution of 25% the original one for Dataset 1. For a resolution of 10%, the accuracy drops to 96.0% . The same consistency can be seen on Dataset 2 until 50% of the original resolution then a drop by 2 points in accuracy is noticed for 25% of the original resolution. The drop further continues with a resolution of 10% the original one. This shows the robustness of the MMLBP algorithm presented in this paper to spatial resolution reduction until a certain percentage depending on the dataset.

Table 5: Accuracy comparison of different spatial resolution

Data-set	Resolution 100%	Resolution 75%	Resolution 50%	Resolution 25%	Resolution 10%
1	99.6 ± 0.4	98.8 ± 0.4	99.4 ± 0.4	98.7 ± 0.4	96.3 ± 0.4
2	99.5 ± 0.3	99.8 ± 0.3	99.5 ± 0.3	97.6 ± 0.3	96.0 ± 0.4

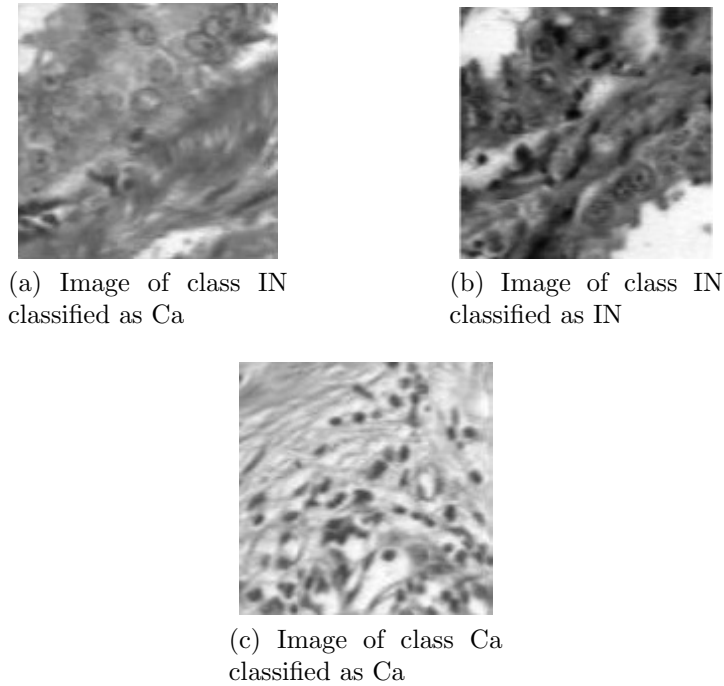


Figure 11: Example of correctly classified and misclassified samples from dataset 1.

435 6.5. Comparison to Existing Algorithms

436 Table 6 depicts the performance accuracy obtained when comparing the
 437 proposed algorithm against some existing methods in the literature. [24]’s
 438 algorithm is tested on both the Texas and the Prostate datasets. The au-
 439 thors of [16] report a 98.9 % accuracy on the Texas dataset. Masood *et al.*’s
 440 algorithm is evaluated using the Texas and Prostate datasets using a multi-
 441 class classifier instead of the authors’ binary classifier [15]. As can be seen
 442 in Table 6, the proposed method outperforms these three other algorithms
 443 in terms of accuracy. Tahir *et al.*’s algorithm is evaluated using the prostate
 444 dataset as reported by the authors who achieved a 98.9 % accuracy. The
 445 proposed algorithm is implemented on the same dataset and the results of
 446 99.5 % accuracy clearly show that the proposed technique outperforms [23]’s
 447 algorithm.

448 6.6. Extension to the Infrared Spectrum

449 The algorithm is first evaluated on the visible and near infrared ranges
 450 separately on Dataset 3. Once this done, it is evaluated on a combined

Table 6: Accuracy comparison to literature methods

Data- set	Khelifi <i>et al.</i> [24] (%)	Tahir <i>et al.</i> [23] (%)	Chaddad <i>et al.</i> [16] (%)	Masood <i>et al.</i> [15] (%)	Proposed algorithm: Stacked MMLBP + GLCM (%)
1	89.9	n/a	98.9 ± 0.1	86.3 ± 0.3	99.6 ± 0.4
2	75.6	98.9	n/a	85.1 ± 2.0	99.5 ± 0.3

dataset including both the Vis and IR data by fusing the accuracy results at a score level using the stacking technique discussed in Section 3.2. Table 7 proves that using both the visible and infrared ranges of the light spectrum improves slightly the results. On the Qatar dataset, the proposed algorithm scores 99.2 % when using only the bands representing the wavelengths in the visible spectrum; this same algorithm scores 99.5 % when using the wavelengths from the infrared as well as the visible range. One can notice that the IR alone does not perform as well as the Vis spectrum with this algorithm but it adds different information and helps improving the accuracy when combined.

Table 7: Accuracy of proposed algorithm on Qatar dataset

Dataset	Accuracy
Dataset 3 Vis	99.2 ± 0.1
Dataset 3 IR	96.2 ± 0.5
Dataset 3 Vis+IR	99.5 ± 0.1

7. Conclusion

Multispectral texture features form an attractive method for extracting information from histologic images of colorectal or prostate tumor tissue for classification purposes. This paper proposed a MMLBP feature combined with GLCM using a stacked generalization for feature fusion at the score level for classification. The proposed method showed that this technique gives better results than similar and existing ones available in the literature attaining a classification accuracy above 99 % on all the datasets tested. This

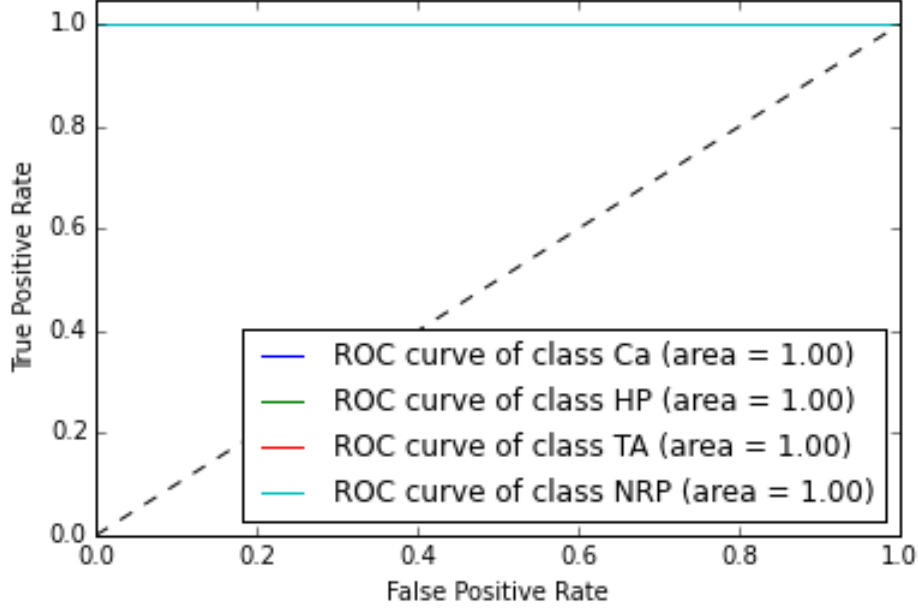


Figure 12: ROC for the proposed algorithm for dataset 3

study also showed that results can be improved when combining both infrared and visible information extracted from tissue samples.

Future work will focus on investigating the use of morphological features in order to improve the result. They will be easily combined to the texture features at decision level for classification thanks to the stacked generalization technique used here.

Appendix A. GLCM Texture Features

The GLCM texture features [9] are calculated from the GLCM extracted from the different layers of the multispectral [4] image where each layer represents the tissue response to a different wavelength. This GLCM matrix reflects how often a pixel with the intensity value I occurs in a specific spatial relationship (r, θ) to a pixel j . Four different spatial relationships are computed: $r = 1$ and $\theta = 0, 45, 90, 135$.

$$GLCM_{i,j,\lambda} = \sum_{p=1}^n \sum_{q=1}^m \begin{cases} 1, & \text{if } \begin{cases} I(p, q, \lambda) = i \\ I(p + \Delta x, q + \Delta y, \lambda) = j \end{cases} \\ 0, & \text{otherwise} \end{cases}$$

482 The following GLCM features are computed from the normalized GLCM
483 matrices $p_{r,\theta}(i, j, \lambda)$ of the image:

- Energy:

$$\sum_{i,j} p(i, j, \lambda)^2$$

- Contrast:

$$\sum_{i,j} |i - j|^2 p(i, j, \lambda)$$

- Homogeneity:

$$\sum_{i,j} \frac{p(i, j, \lambda)}{1 + |i - j|}$$

- Correlation:

$$\sum_{i,j} \frac{(\mu_i - i)(\mu_j - j)}{\sigma_i \sigma_j} p(i, j, \lambda)$$

484 For each multispectral image, the GLCM features are calculated on each
485 GLCM from each layer and concatenated into a large vector of size $4 * \text{number of multispectral layers}$. The features are then rescaled and nor-
486 malized to fit in the interval $[0, 1]$ using the following equation:
487

$$x = \frac{x - \min(x)}{\max(x) - \min(x)} \quad (\text{A.1})$$

488 Where x is a vector representing the feature to be normalized, x' is the
489 normalized feature, $\max(x)$ and $\min(x)$ are respectively the maximum and
490 minimum values of x .

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