Dynamic Low-Level Context for the Detection of Mild Traumatic Brain Injury

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Abstract— Mild traumatic brain injury (mTBI) appears as low contrast lesions in magnetic resonance (MR) imaging. Standard automated detection approaches cannot detect the subtle changes caused by the lesions. The use of context has become integral for the detection of low contrast objects in images. Context is any information that can be used for object detection but is not directly due to the physical appearance of an object in an image. In this paper new low level static and dynamic context features are proposed and integrated into a discriminative voxel level classifier to improve the detection of mTBI lesions. Visual features, including multiple texture measures, are used to give an initial estimate of a lesion. From the initial estimate novel proximity and directional distance contextual features are calculated and used as features for another classifier. This feature takes advantage of spatial information given by the initial lesion estimate using only the visual features. Dynamic context is captured by the proposed posterior marginal edge distance context feature, which measures the distance from a hard estimate of the lesion at a previous time point. The approach is validated on a temporal mTBI rat model dataset and shown to have improved dice score and convergence compared to other state-of-the-art approaches. Analysis of feature importance and versatility of the approach on other datasets are also provided.

Index Terms— Magnetic Resonance Imaging, Traumatic Brain injury, Low Contrast, Dynamic Context.

I. INTRODUCTION

Awareness of mild traumatic brain injury (mTBI) has increased dramatically in recent years. Causes of mTBI include sports injuries, automobile accidents, blast injuries in the military, and falls in the workplace [1]. The long term effects of mTBI are just being recognized, leading to the need for quantitative techniques to characterize and measure the

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injured tissue.

Clinical evaluation of mTBI has been too qualitative by relying on the Glasgow Coma Scale, to assess loss of consciousness, loss of memory, alteration in mental status, and focal neurological deficits. When imaging is used to assist in diagnosis, MR imaging or computed tomography (CT), a quantitative measurement of the size and location of injury is typically obtained. The primary focus of imaging is only to assess for the presence of a hematoma [1]. Various clinical studies have used manual quantitative analysis to evaluate TBI in MRI [2-4]. These studies identified the size and location of the lesions and linked them to long term neurological effects. Manual analysis has some major downfalls; hours per scan of ROI extraction, a trained operator it requires and difficulty in interpreting multiple modalities. Inter and intra operator error can become significant in low contrast images. Inter operator error has been shown to be significant even in higher contrast tumors [5]. However, manual segmentation is considered the "gold standard" and is used as the ground-truth.

Some computational approaches have been proposed for quantifying lesions in moderate to severe TBI [6], which on MRI have high contrast, but these have been unsuccessful when attempting to evaluate the subtle MR signature of mTBI. Lesions caused by mTBI appear as small low contrast regions (Figure 1) in both T2 weighted images and T2 maps. The T2 values within these lesions often fall within the range of

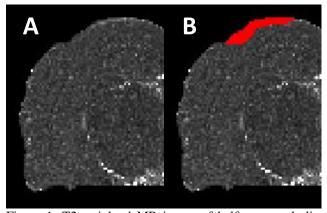


Figure 1: T2 weighted MR image of half a coronal slice from the rat model dataset. A) Original T2 weighted image. B) Manual detection of the mTBI lesion (highlighted in red). This illustrates the low contrast appearance of mTBI lesions. The manual lesions have been verified by histology in [13].

normal tissue values. Therefore, the T2 value of a voxel cannot be used by itself. In [7] it was shown that there are significant texture changes in brain tissues as a result of mTBI, which provides a measure of the underlying structure of the tissue. Therefore, our proposed approach uses multiple texture measures to improve the discriminative ability of mTBI detection from MR images.

There has been research into advanced MR techniques for detecting mTBI, mainly susceptibility weighted imaging (SWI) [8-10] and diffusion tensor imaging (DTI) [10-12]. SWI utilizes the phase to estimate the magnetic susceptibility in tissue. It has been shown to be more sensitive to blood than CT. However, SWI tends to overestimate a lesion and can give too sharp a contrast leading to hard determination of lowcontrast brain structures [9]. DTI measures the "flow" of water in the brain, and has been used to estimate white matter (WM) tracts. Signification changes in WM tracts have been found in patients with mTBI, but it requires multiple imaging time points or a baseline image [14]. It becomes difficult to evaluate it due to inter patient variations in tractography. This rules out DTI for voxel based classification. We utilize T2 weighted imaging, a traditional imaging technique as described above.

A measure of contrast within the lesion can be obtained by calculating the contrast-to-noise ratio (CNR). The manually segmented regions are used to calculate the CNR, and a CNR of 6 on average is obtained. Rajin et al. [15] made noise measurement from MRI to estimate the noise variance.

To estimate the mTBI lesion we use a discriminative approach comprised of voxel level classification utilizing visual and contextual features. Both static and dynamic contextual features are proposed. Static contextual features, proximity and directional, are calculated from local neighborhood information of the posterior marginal estimated by a preceding classifier. The dynamic context (posterior marginal edge distance) is calculated from the maximum posterior marginal of the cascade of classifier of a previous time point and then used to estimate the current lesion. Adding these contextual features overcomes the low contrast nature of the lesions.

The paper is organized as follows: Section 2 presents the related work and the contributions. Section 3 discusses the technical approach. Section 4 describes the experiments and finally Section 5 provides the conclusions.

II. RELATED WORK AND CONTRIBUTIONS

A. Related Work

Context has been an active are of research in computer vision and can be used to overcome low contrast detection problems. Context is defined [16] as, "any information that might be relevant to object detection, categorization and classification tasks, but not directly due to the physical appearance of the object, as perceived by the image acquisition system." Context types that have been described include: local pixels, 2D scene gist, 3D geometric, semantic, photogrammetric, illumination, weather, geographic, temporal,

and cultural [17]. Context can be split into two types local and global: a) Local context includes spatial relationships learned at the pixel/region level. It is used in conditional random fields [18] and auto-context [19], and b) Global context can be thought of as an estimation of the spatial location of an object. In this paper we exploit low (voxel) level context. Our previous approach [20] uses a contextually driven generative model to estimate the lesion location.

As a starting point algorithms for Multiple Sclerosis (MS) are examined due to similar low contrast lesions being present in MR images. The approaches for detecting MS lesions [21-22] use texture features and take advantage of the knowledge that MS occurs in a specific tissue type, which is consistently located in similar anatomical regions. This is a type of context (anatomical context) that can be exploited easily for MS, but mTBI lesions do not have this advantage since mTBI can occur in different tissue types.

Recently [19] proposed autocontext is a way to model context at the voxel level. The main premise is to estimate an object with a discriminative classifier and use a sampling of the estimated posterior probability as an additional feature to a subsequent classifier. It is able to take information from far away (across the brain) compared to other methods like conditional random fields (CRF) [18] which are local. The context features in this case are a sparse sampling of a distant neighborhood around every pixel. This can lead to overfitting due to the very specific locations and this can be seen in one of the examples in [19]. Other methods that exploit this type of context include [23], which has similar pitfalls as

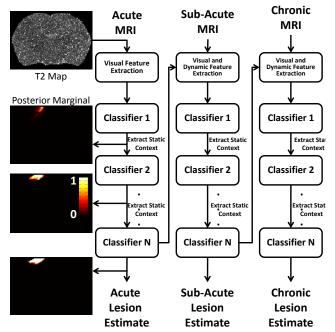


Figure 2: Overview of the proposed system. Context information is sent from one classifier to the next. After each classifier the static contextual features are extracted. The output of the Nth classifier is the final estimate of the lesion for the given time point. The dynamic contextual features are estimated from the final posterior marginal of the preceding time point.

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autocontext. These methods have been shown to segment rigidly positioned objects well, such as organs or body parts. In this paper we adopted the idea of cascading classifiers, but developed features that are more generalizable and integrate temporal information, which has not been addressed in [19].

B. Contributions

The contributions of our paper are: 1) development of three new contextual features to be used with a cascade of classifiers. These features include a proximity feature, a directional feature, and a maximum a posteriori edge distance feature. 2) Use of a temporal sequence of MR images to provide context from a previous time point, capturing the dynamics of the injury progression automatically. 3) Analysis of multiple contextual feature configurations on a rat controlled cortical impact (CCI) mTBI model dataset.

III. TECHNICAL APPROACH

A. System Overview

A discriminative approach is taken where classifiers are used to directly estimate the posterior probability of lesion and normal appearing brain matter (NABM) voxels (Note the ground-truth is obtained from segmentation of the mTBI lesion by a domain expert). The discriminative approach performs well when there is a large amount of training data and it can be used for a complex decision space. A voxel level classifier has a large amount of data considering the 3D nature of MR images. The appearance of lesions in MRIs can be very complex, which leads to a complex decision space.

Here a cascade of classifiers is used for estimating the detected lesion at each time point (shown in Figure 2), where information from a previous time point is propagated forward. The first classifier in the cascade estimates the lesion using only the visual features. Then context features are formed from the posterior probability map estimated by the classifier. These features are recalculated for each iteration in the process, for a given number of classifiers shown as a column of classifier for a single time point in Figure 2. Spatial information is propagated by the contextual features leading to improved classification. This process was introduced by [19]

for static images. The contextual features used by [19] were point samplings of the posterior estimates. Their features demonstrated good segmentation performance on objects that were rigid in shape, when that shape was distorted their performance faltered. We propose two new features that generalize the static contextual information so it will work with amorphous objects.

Dynamic contextual features are calculated from the final classifier at a single time point. These features also in our approach are used by classifier at the subsequent time point, and they make use of spatial and lesion growth information. Note that [19] proposed using spatio-temporal volumes with their basic point features in the higher dimensional space. Their approach would require extreme amounts of data and the entire sequence to be known before segmentation. Our approach only considers pairs of brain volumes at a time, which allows for estimation at every time point.

B. Visual Features

Due to the low contrast nature of the unimodal MR images and the mild nature of the TBI, texture features are used to increase the discrimination in our approach. Four types of texture features are used: uniform local binary pattern (ULBP) local histograms [24] in the coronal plane (59 features), local statistics (mean, variance, skewness, kurtosis) of a Gabor filter bank [25] with 8 orientations and 4 scales in the coronal plane (128 features), basic histogram of oriented gradients in the coronal plane [26] (9 features), and local neighborhood statistical features (mean, variance, skewness, kurtosis, range, entropy, gradient magnitude xyz) (9 features). This gives a total of 205 visual features. This wide variety of features provides many different characteristics without being too specific (i.e., they will generalize well). The size of the local neighborhood used is 5x5 voxels [33], where the feature size is chosen to be the one that maximizes the Bhattacharyya distance between the two classes and gives a bound on the Bayesian error [27]. Examples of the visual features are shown in Figure 3.

C. Static Contextual Features

The contextual features come from the posterior probability

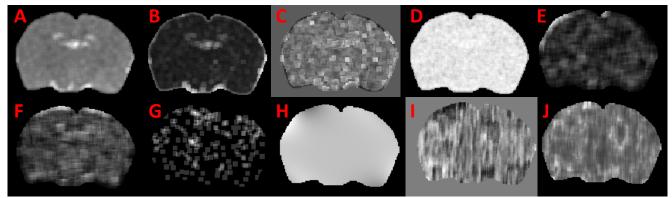


Figure 3: Examples of the visual features on coronal slices. The features from left to right, top to bottom: local mean (A), variance (B), skewness (C), kurtosis (D), a bin from the histogram of oriented gradients (E-F), a bin from the LBP histogram (G), and example gabor features (H-J).

estimated by an already learned classifier. Previous approaches [13] have directly sampled a dense neighborhood around an observed voxel, making each location a potential feature. This method can lead to large feature sizes and can cause overfitting due to the specific locations that are learned. In this paper, two new static features are proposed to overcome this problem. One incorporates a sense of the surrounding without a known direction, while the other gives a general sense of direction.

The first feature, shown in Figure 4A, gives the average posterior probability at various distances around the observed voxel. This can be thought of as a proximity feature where: what is a close, medium and far away in distance are estimated. The distance function used here is the Manhattan distance allowing for a cuboidal region. These features are directionally invariant and can lead to better generalization since they describe a general area. By having a nesting of boxes the integral image can be utilized for quick computation of the features. In 3D only 8 points are needed to find the average of a cuboidal region, using integral images [28]. Equation 1 provides these features, where f_{xyz} is the proximity feature, R_{1xyz} , R_{2xyz} are square neighborhoods around the voxel at xyz and size(*) is the size of the bounding box.

$$f_{xyz} = \left(\sum R_{1xyz} - \sum R_{2xyz}\right) \left(\frac{1}{size(R_{1xyz}) - size(R_{2xyz})}\right) \quad (1)$$

Directional information is important for classification since the objects are rigidly registered to a naïve animal. The second contextual feature describes the posterior probability in various directions from the observed voxel. Rays are sampled at various distance ranges and angles from the observed voxel (see Figure 4B). From the distance ranges along the rays the mean is calculated. This gives a refined sense of the surrounding. An example would be what is close and above the observed voxel. The integral image is also used to calculate these features. Both features can be computed at coarse or fine distance bins without a significant increase in computational time. Equation (1) can also be used for the

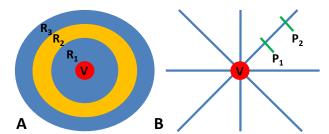


Figure 4: A) Illustration of the proximity feature. V is the observed voxel and the feature is the average probability of the regions (R_1 , R_2 , R_3). B) Illustration of the distance features. V is the observed voxel and an example feature is the average probability between P_1 and P_2 along the 45° ray.

direction context features, the shape of the features are changed to be that of Figure 4B. 3The features are rectangles with width one and the 45 degree features can be estimated using the rotated image.

D. Dynamic Contextual Feature

We propose the posterior marginal edge distance (*PMED*) feature is the distance a voxel is from the perimeter of objects of a class found by the maximum posterior marginal (*MPM*) estimate. To create this feature first the *MPM* at a voxel is obtained from the output of a classifier. This gives a binary image for each class. The distance transform is applied to the image and the inverse image and the feature is given by equation 2.

$$PMED = d(MPM) - d(\sim MPM) \tag{2}$$

$$MPM = \mathop{argmax}_{c} p(\omega = c|F)$$
 (3)

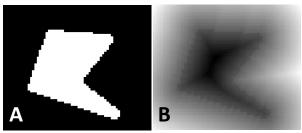


Figure 5: A) Example MPM estimate. B) Corresponding PMED feature. Note that the values become more negative towards the center of the object and more positive farther away from the object.

Here d(*) is the Euclidean distance transform. This gives an image that is increasing as the voxels become farther away from the edge and smaller (more negative) as the voxels get further into the object. In eq. (3) ω is the estimated class, c is a specific class (lesion or normal brain in our case), and F is the feature set at a given voxel (see Figure 5).

E. Classifier Cascade Selection

The base classifier in the cascade of classifiers has to be able to deal with a large number of features, a complex decision space and be able to give a posterior estimate. The primary classifier being used is adaboost [29] with small decision trees as base classifiers. Using small trees as a weak classifier (h(*)) allows for inherent feature selection, meaning erroneous features are disregarded. It is also not sensitive to feature normalization. In each iteration (t), the best classifier is selected and weighted with α . During the training process a cost matrix (4) is used where the top row represents the cost of normal appearing brain matter (NABM) and the bottom row the cost of Lesion voxels, such that the priors are offset to be even. This is done to account for the large disparity, in the number of training samples between the classes. Algorithm 1 describes the flow of adaboost.

$$Cost\ Matrix = \begin{bmatrix} 0 & 1\\ \frac{\sum NABM\ voxels}{\sum Lesion\ voxels} & 0 \end{bmatrix} \tag{4}$$

It has been shown that the posterior marginal can be estimated using logistic regression [30] given by equation 5.

$$p(\omega_{xyz} = c | f_{xyz}) = \frac{e^{H_c(f_{xyz})}}{\sum_{c=1}^{C} e^{H_c(f_{xyz})}}$$
(5)

$$H_c(f_{xyz}) = \sum_{t=1}^{T} \alpha_t h_t(f_{xyz})$$
(6)
Other classifiers such as the popular support vector machine

$$H_c(f_{xyz}) = \sum_{t=1}^{T} \alpha_t h_t(f_{xyz})$$
 (6)

(SVM) approach are also able to estimate the posterior marginal [31]. SVM is a strong classifier, but it does not have inherent feature selection and is sensitive to feature normalization. SVM tends to perform well with a small number of strong features. An example of the iterative training process for a single time point is shown in Figure 6. Notice that the estimate improves the greatest between classifier 1 (Figure 6B) and classifier 2 (Figure 6C). During training each successive classifier will have a performance greater than or equal to its predecessor, since the posterior marginal is one of

ALGORITHM I ADABOOST

ADABOUST	
1:	Input(x,y) where x is the set of all training features and features and y is the class label.
2:	T = number of weak classifiers
3:	Initialize $D_1 = 1/m$ where D is the weight of each training example and m is the number of training examples.
4:	For $t = 1$ to T
5:	$h_t = argmax_{h_t \in H} 0.5 - \epsilon_t // Base Classifier$
	where $\varepsilon_t = \sum D * \sum Cost Matrix * (y \neq h_t(x)) / Error$
6:	$\alpha_t = \frac{1}{2} ln \left(\frac{1 - \varepsilon_t}{\varepsilon_t} \right) // \text{ Classifier Weight}$
7:	$D_{t+1} = \frac{D_t \cdot \exp(-\alpha_t y h_t(x))}{Z} // \text{Reweight the examples}$
	Where Z is a normalization term.
8:	Exit if $ 0.5 - \epsilon_t < threshold$ //exit if error is small

the features.

IV. EXPERIMENTAL RESULTS

A. Contribution of Features in a Temporal mTBI Dataset

Our temporal data set uses Sprague Dawley rats subjected to a single impact controlled cortical impact (CCI) as an animal model of mTBI [13, 32]. The CCI model results in a focal injury. Briefly, a craniectomy (5mm) was performed over the right hemisphere (3mm lateral, 3mm posterior to bregma) where a mild CCI was induced using an electromagnetically driven piston (4mm diameter, 0.5mm depth, 6.0 m/s). The incision made for the craniectomy is sutured after the injury is induced.

The animals were imaged in vivo at 3 time points post injury: acute (1st day), sub-acute (8th day), and chronic (14th day). There are a total of 6 sequences, each with 3 time points. MRI data were acquired using a Bruker Advance 4.7T for T2 weighted images (T2WI; TR/TE/FA=3453 ms/20 ms/20°, 25x1 mm slices) with a 256x256 matrix and 3cm field of

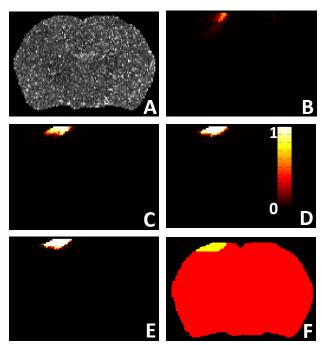


Figure 6: Example probability maps after each classifier. A) T2 map. B-E) Classifier output probability map for the training of classifier 1-4 respectively. (F) Manual Segmentation where yellow denotes the lesion. This shows the convergence of the algorithm to the manually segmented injury.

view. Each voxel has a dimension of 0.12x0.12x1mm. The T2 weighted images were converted to T2 maps. ROIs were manually segmented using Cheshire image processing software (Hayden Image/Processing Group, Waltham, MA) and included the right and left hemispheres and injured tissue volumes that were defined as abnormal (hyper/hypo-intense) signal intensities within the cortex with the remaining tissues designated as NABM. The injuries have been verified with histology in [32] at the end points of experiments. A direct comparison to the results of histology is not possible due to the deformations that occur in the histological staining process.

We examine the effect of the proposed features and effect of the dynamic information. For the training/testing split leave-one-out validation is used where a whole sequence is left out (resulting in 6 cross-folds). Three temporally consecutive volumes were left for testing and the rest were used for training. The parameters used were: 300 weak learners, learning rate 1, and 4 cascaded classifiers. Three approaches were tested: the original autocontext features [19], the proposed approach with only the proposed static features, and the proposed approach with the static and dynamic features.

To evaluate the segmentation results of the proposed approach the dice coefficient is used, given by equation (7),

$$Dice = \frac{2TP}{(TP + FP) + (TP + FN)} \tag{7}$$

where TP is true positive, FP is false positive, and FN is false

negative. The dice coefficient gives the ratio of the intersection between the detected object and the target object to the size of the objects. The Dice gives a better idea of the intersection between the two objects because it does not consider the true negatives, which will be large with the unbalanced class size. The output of the classifier at each time point (acute, sub-acute, and chronic) shown in Figure 2, is thresholded to get a hard classification.

From Figure 7 it is clear that the proposed dynamic approach outperforms the other methods. This shows it is important to use the dynamic information. The original autocontext tends to over fit due to the specific locations the features represent. The same features locations proposed in [19] were used for the original autocontext testing. During the training phase it obtains a dice score above 0.9, but it does not generalize well to the testing data. From the testing, our proposed static features give a good generalization compared to the original autocontext. The output from the original autocontext approach is patchy with many false negatives leading to an under-estimation of the injury. This is due to the fact that it looks at specific locations for the context features.

The proposed approach has a very flat dice curve, so it is not sensitive to a chosen threshold on the output probability map. This makes the selection of a threshold less critical. From the qualitative results (Figure 8) it is clear that the results of proposed approach work on small to medium size lesions. The qualitative results also show that false positives are only close to the majority of the lesion mass without having erroneous spatial outliers. It is also apparent that the approach has some difficulties at the edges of the lesions. This could be rectified by using shape constraint on the thresholded probability map.

In [20] a similar dataset was used for analysis except repeated contralateral injuries were also considered. The average dice score using a high level contextual model was 0.35 and 0.07 using probabilistic SVM alone. This is similar to our dice score using the proposed static features with an average dice score of 0.36 and it far exceeds the results of a probabilistic SVM. The dynamic features increase the dice up to 0.41. The method proposed here does not use any extra patient information as it has been done in [20].

B. Convergence Rate of the Features

We evaluate the strength of the individual features. The same dataset from section IV.A is used in this section. To determine the strength of the features the rate of convergence of the cost function during training is observed.

In Figure 9 the training cost after each tree during the training of adaboost is given. Both the autocontext and the proposed static features have the same results from the initial classifier utilizing only visual features. This is due to the proposed static features being able to generalize in the surrounding area better.

To understand the convergence of each classifier in the cascade one must observe the output training cost after adding each tree during the training of adaboost. In Figure 10 the convergence of the training using only the static features is

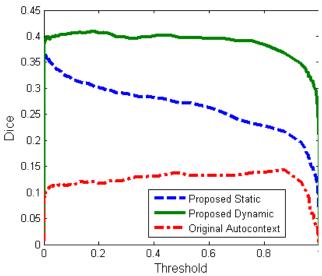


Figure 7: Dice coefficient after thresholding the posterior probability map at the end of each cascade of classifier (i.e. at each time point). This is the average of all the tests in the leave one out validation.

observed. Convergence using only the visual features takes a while compared to with the static contextual features. After the second classifier, the improvement over the number of trees is less pronounced. Not much information is gained after the second classifier.

Also in Figure 10 we can observe the convergence of the classifier using the dynamic and static context features. It is apparent that the convergence is very fast for Classifier 1 when the dynamic context features are used, compared to the case when no dynamic information is used in the first iteration. Information is still gained by the static features but at a slower rate.

C. Feature Importance Evaluation

An important question to ask when using adaboost is, "which features are being chosen and when?" Here those questions are answered by looking at the percent of features chosen throughout the adaboost training process. In the case of the static features alone, Figure 11 shows first 50 trees are almost exclusively context features. This shows that the contextual features hold the most information for discrimination. The visual features start being used after the information from the contextual features is nearly all used. In the end 40-50% of all the features are contextual.

A similar pattern occurs when the dynamic features are used (shown in Figure 11). The contextual features in Classifier 1 are used early on in the learning process, and then the visual features almost exclusively take over. In Classifier 3 nearly 60% of the features are chosen as contextual features. This shows that the dynamic information plays a vital role in the classification.

D. Static Feature performance on the BRATS dataset

To show the versatility of the proposed approach the static features were tested on a brain tumor dataset. The dataset is from the MICCAI 2012 Multimodal brain tumor segmentation

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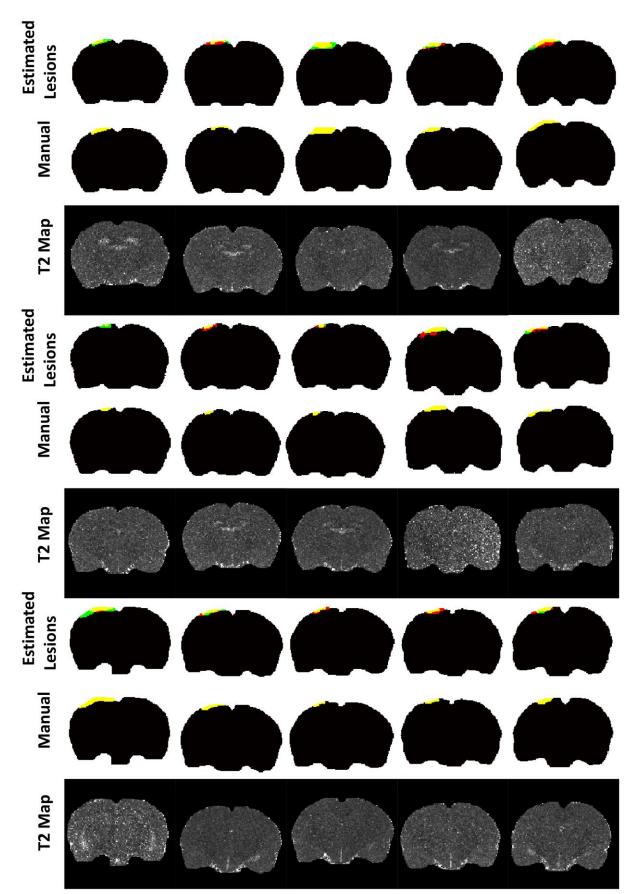


Figure 8: Qualitative results of the proposed approach using dynamic and static contextual features. Each coronal slice is from a separate volume. Color code: yellow = true positive, black = true negative, green = false negative, red = false positive.

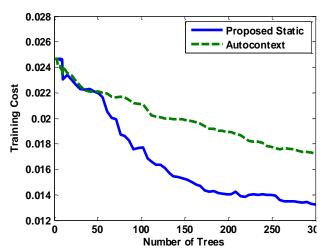


Figure 9: Training cost after each tree for the proposed static features versus the original autocontext. This is the training cost of the second classifier i.e. the input to both methods is the results after using visual features only. Lower cost is better.

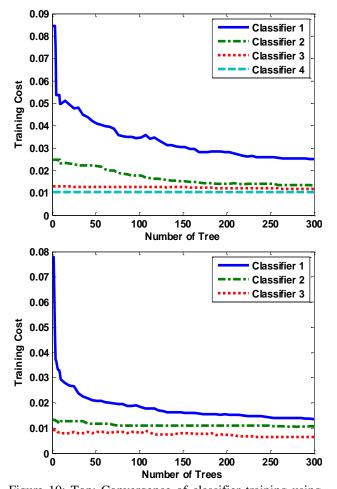


Figure 10: Top: Convergence of classifier training using only the static contextual features. Bottom: Convergence of classifier training using both dynamic and static contextual features. Lower cost is better.

(BraTS) challenge. Here a subset of the data was used that included 15 High grade glioma patients. The volumes included

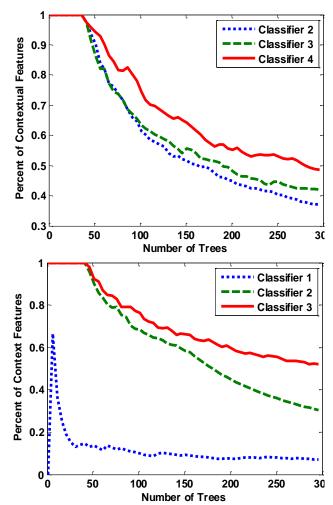


Figure 11: Top: The percentage of the static contextual features chosen versus the current number of trees. Bottom: The percentage of dynamic and static contextual features chosen up to a number of trees. Note in Classifier 1 only the dynamic contextual features are used.

the following MR modalities, T1, post-Gadolinium T1, T2, and FLAIR. Each modality has been co-registered, skull-stripped and resampled to be 1mm isometric voxels. A mode shift was used to normalize the histogram across subjects. For this testing the edema and core tumor are considered to be a single class. The visual features used were the individual modalities, the same local neighborhood statistics as the mTBI dataset on the FLAIR channel, and LBP on the FLAIR channel. Since there is a single time point database only the static contextual features are used. Testing was done using 5 fold-cross validation.

For this testing an average dice score of 0.71 was obtained. The best scores reported during the BraTS challenge were between 0.7 - 0.8, for the high grade glioma cases. Note that the other approaches [33, 34] used symmetry features. These features can be directly used by the proposed approach. This is another type of context that can be exploited. Another feature used in the challenge were long range context features [35]. These features sample the value around an observed voxel. The parameter space for these features is extremely large and

requires random sampling of the space via decision forests [36]. The other features used by the winners of the challenge [35] were based on generative modeling of the tissue types. These features can also directly be used by the approach proposed here.

V. CONCLUSIONS

A fully automated method for detecting mTBI lesions that integrates low-level static and dynamic context is proposed. Three novel features are proposed that describe the posterior probability of classifier outputs in a cascade. These features were shown to have good qualitative and quantitative results on an mTBI dataset. The proposed approach outperformed the original autocontext [19], by being able to generalize and integrate dynamic information. The context features were shown to be the most important features in the training process. Adding the dynamic context features lead to fast convergence during training leading a smaller number of trees being needed. The generality and flexibility of the proposed approach allows it to be applied to other brain lesion problems.

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