

Image Similarity and Asymmetry to Improve Computer-Aided Detection of Breast Cancer

Dave Tahmoush^{1,2} and Hanan Samet¹

- ¹ Computer Science Department, Center for Automation Research, Institute for Advanced Computer Science, University of Maryland, College Park, USA, tahmoush@cs.umd.edu, hjs@umiacs.umd.edu,
² Applied Physics Laboratory, Johns Hopkins University, Laurel, Maryland, USA

Abstract. An improved image similarity method is introduced to recognize breast cancer, and it is incorporated into a computer-aided breast cancer detection system through Bayes Theorem. Radiologists can use the differences between the left and right breasts, or asymmetry, in mammograms to help detect certain malignant breast cancers. Image similarity is used to determine asymmetry using a contextual and then a spatial comparison. The mammograms are filtered to find the most contextually significant points, and then the resulting point set is analyzed for spatial similarity. We develop the analysis through a combination of modeling and supervised learning of model parameters. This process correctly classifies mammograms 84% of the time, and significantly improves the accuracy of a computer-aided breast cancer detection system by 71%.

1 Introduction

Breast cancer remains a leading cause of cancer deaths among women in many parts of the world. In the United States alone, over forty thousand women die of the disease each year [1]. Mammography is currently the most effective method for early detection of breast cancer [2]. For two-thirds of the women whose initial diagnosis of their mammogram is negative but who actually have breast cancer, the cancer is evident upon a second diagnosis of their mammogram [2]. Computer-aided detection (CAD) of mammograms could be used to avoid these missed diagnoses, and has been shown to increase the number of cancers detected by more than nineteen percent [3]. Measuring asymmetry, which consists of a comparison of the left and right breast images [4], is a technique that could be used to improve the accuracy of CAD. An automated prescreening system only classifies a mammogram as either normal or suspicious, while CAD picks out specific points as cancerous [5]. One of the most challenging problems with prescreening is the lack of sensitive algorithms for the detection of asymmetry [6]. This paper presents a simple and effective algorithm for the detection of asymmetry and extensions to improve upon it. We improve on our earlier results [7] and incorporate image similarity into a CAD system.

Mammograms are an excellent candidate for image similarity techniques to be effective because there are images of both the left and right breasts, which should

be similar if there is no cancer present. Image similarity has been often utilized for content-based image retrieval (CBIR) from image databases [8–11]. Both contextual and spatial comparisons are used [8]. Medical image databases have also used image similarity, from rule-based systems for chest radiographs [12] to anatomical structure matching for 3D MR images [13]. However, the focus is often on the non-cancerous structures, while it is the cancerous structures that are of principle interest here. In this paper we combine the image similarity concept of contextual then spatial comparison to the problem of detecting breast cancer in mammograms.

The majority of work on CAD analysis of mammograms has focused on determining the contextual similarity to cancer, finding abnormalities in a local area of a single image [14, 15]. This paper focuses on combining this with a spatial comparison in order to complete an image similarity measure. The majority of work has used methods ranging from filters to wavelets to learning techniques, but a detailed discussion of various imaging techniques is beyond the scope of this paper. Problems arise in using filter methods [14] because of the range of sizes and morphologies for breast cancer, as well as the difficulty in differentiating cancerous from non-cancerous structures. The size range problem has been addressed by using multi-scale models [15]. Similar issues affect wavelet methods, although their use has led to reported good results [16] with the size range issue being improved through the use of a wavelet pyramid [17]. Learning techniques have included support vector machines [18] and neural networks [16].

Detecting breast cancer is challenging because the cancerous structures have many features in common with normal breast tissue. This means that a high number of false positives or false negatives are possible. Asymmetry can be used to help reduce the number of false positives so that true positives are more obvious. Previous work utilizing asymmetry has used wavelets or structural clues to detect asymmetry with correct results as often as 77% of the time [4, 19]. Additional work has focused on bilateral or temporal subtraction, which is the attempt to subtract one breast image from the other [20, 21]. This approach is good because it does try to utilize the multiple images taken with the same machine by the same technician and analyzed using the same process in an effort to reduce the systematic differences that can be introduced. However, bilateral subtraction is hampered by the necessity of exact registration and natural asymmetry of the breasts. We introduce a measure of asymmetry that is more approximate in nature and seems more robust to the large amount of noise in the data, using learning to determine a highly constrained number of model parameters. Minimizing the number of parameters that are learned makes the model less subject to overfitting the noise in the data at the possible expense of accuracy.

Comparing multiple mammograms using learning techniques has been shown to be effective in CBIR [10, 22]. Our application lends itself well to supervised learning because the data set has already been screened for cancer and thus classified by expert radiologists. However, care must be taken since the expert classification is known not to be perfect [2].

We believe that developing ways to better utilize asymmetry is consistent with a philosophy of trying to use methods that can capture measures deemed important by doctors thereby building upon their knowledge base, instead of trying to supplant it. However, measuring asymmetry means comparing multiple images, and thus it is a more complicated process.

In order to more fully understand the effectiveness of incorporating asymmetry into CAD systems, we utilize Bayes Theorem as a simplified method for combining a global measurement of asymmetry with the local measure of the probability of having a cancer. Significant improvement is shown over the system without asymmetry.

The rest of this paper is organized as follows. Section 2 presents our method for measuring asymmetry between the mammogram images and incorporating the measure into a CAD system. Section 3 discusses the evaluation of the performance of the measure and compares the results with other work. Section 4 describes future work, while Section 5 discusses the conclusions that can be drawn from this work.

2 Asymmetry Measurement And CAD

Our work utilizes filtering followed by spatial analysis to determine an overall measure of similarity by combining the contextual similarity of the filtering with the spatial similarity of the analysis. This can be a useful measure for prescreening mammograms since only an overall determination is required. It can also be incorporated into CAD as we demonstrate using Bayesian statistics. A secondary goal of our work is to determine the importance of similarity or asymmetry in the computer analysis of mammograms.

Our analysis starts with filtering to find the contextually similar suspicious points that could be cancers in the mammograms. The filtering step is the same as we used in [7]. This yields a set of potential detection sites that can be analyzed for asymmetry. Although it may not be the optimal choice of either filtering or ranking, the spatial analysis that we used can be applied to any technique that can rank the suspiciousness of areas. The number of points returned by the filtering step is one of the variables that can be adjusted to optimize the analysis. Alternatively, we can also make use of a threshold on the suspiciousness value instead of taking the top few. However, we chose to take the top few in order to be insensitive to image processing choices that might bias the analysis.

The analysis for similarity or asymmetry that we used does a comparison of the values of the sets of suspicious points. Two separators are learned with a training set of images. A model for a separator in 3D is a plane with the parameter set mx, my, b . Parametric learning is used to determine the best parameters based on the training set. A separator breaks the set of suspicious points in both images into two groups, and the populations in the groups are compared between the images. This is based upon the assumption that the presence of cancer will distort the distribution of suspicious sites, and that the distribution will be very similar from left to right breasts when there is no cancer. This is similar to

comparing histograms whose parameters are learned from a test set. We used learning techniques to determine the optimal structure and parameters for the separator from the data. For this application, the importance of correct classification of the cancerous cases is much more important than the non-cancerous cases. To reflect this, the associated weighting of the cancerous cases was varied, and we evaluate the performance of various weightings.

The two separators break the areas in 3-space into groups labelled $L_1, L_2, L_3, R_1, R_2, R_3$, and each has its associated occupancy of suspicious points. A measure of asymmetry D is then calculated as $D = \sum_j |L_j - R_j|$. A more flexible measure would consider each set of groups as a separate measure with its own threshold. In order to use this measure to classify each case as either cancerous or non-cancerous, a threshold needs to be applied. This threshold is another parameter that is learned.

To incorporate asymmetry into a CAD system, we made use of Bayes Theorem, $P(CancerSite|Asym) = \frac{P(Asym|CancerSite)P(CancerSite)}{P(Asym)}$. The sites where asymmetry is measured are thus given an increased probability of being cancerous, while sites where asymmetry is not measured are given a reduced probability of being cancerous. Since the asymmetry measurement is currently done on an entire case, all of the sites in those cases are affected similarly. The effect can be seen in Figure 1. Using Bayes Theorem to incorporate asymmetry into CAD is shown to work well at low numbers of false positives per image, but the overall performance is still strongly dependent on the effectiveness of the CAD system. The true positive fraction of the asymmetry measurement is essential in order to prevent true positives from having their probabilities diminished, and the false positive fraction is important for improving the effectiveness of the CAD system. At high levels of false positives per image, the incorporation of asymmetry has minimal effect, but this is expected since using Bayes Theorem merely reduces the probability of the false positives and does not eliminate them.

3 Evaluation and Results

The groups of suspicious points in the left and right mediolateral oblique (MLO) mammogram views were compared to evaluate the asymmetry measure. The analysis was done with cases that were normal mammograms and mammograms with malignant spiculated lesions from the Digital Database for Screening Mammography [23]. Spiculated lesions are defined as breast cancers with central areas that are usually irregular and with ill-defined borders. Their sizes vary from a few millimeters to several centimeters in diameter and they are very difficult cancers to detect [17]. The training set had 39 non-cancerous cases and 37 cancerous cases, while the test set had 38 non-cancerous cases and 40 cancerous cases. The data is roughly spread across the density of the breasts and the subtlety of the cancer. The breast density and subtlety were specified by an expert radiologist. The subtlety of the cancer shows how difficult it is to determine that there is cancer. The training data set was used to determine optimal parameters for the

separators used in the classification. These cases indicated that a difference in the groups of one or more suspicious points indicated cancer.

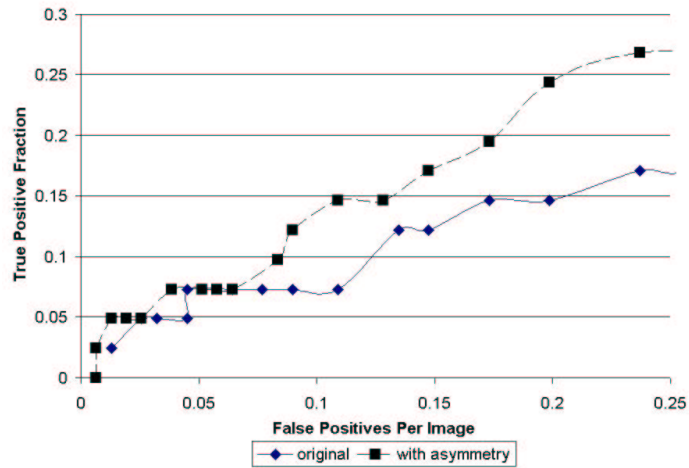


Figure 1: Comparison of the CAD system before and after the addition of asymmetry. The improvement in performance is good at a small number of false positives per image.

Our results are good on all cases of the test set, correctly classifying 84% on the test set. This shows that spatial distribution of suspicious points is changed by the presence of a cancer. However, it is much more important to correctly classify the cancerous cases, and by heavily weighting the importance of the cancerous cases, we correctly classify 97% of the cancerous cases but only 42% of the non-cancerous cases. Neither the subtlety nor the density of the cancer had an effect on the results. The comparison with a commercial system shows that the results are surprisingly good. Correct classification results of 96% of the cancerous cases and 33% of non-cancerous cases are possible using the R2 ImageChecker system [6]. Our method showed correct classification results on 97% of cancerous cases and 42% of the non-cancerous cases. This demonstrates the importance of asymmetry in pre-screening, since using only asymmetry achieves a better performance than a complete commercial system. The inclusion of additional factors other than asymmetry in the method should improve the results. However, the data sets used are different, as the R2 ImageChecker data contains all cancer types and our method has only the difficult to detect spiculated lesions. The R2 ImageChecker data set also had a much higher proportion of non-cancerous mammograms to cancerous cases.

The results on using Bayes Theorem to incorporate asymmetry into CAD were good, increasing the accuracy by up to 71% at a set level of false positives

per image, as is shown in Figure 1. The improvement is most apparent at low levels of false positives. At higher levels of false positives, the effectiveness of the CAD technique dominates because the Bayes technique does not actually remove any false positives, it merely reduces their probability of being cancerous.

Our method makes use of a spatial analysis of the suspicious points, counting the number of suspicious points in the groups created by the separator. Its success is an encouraging sign for the investigation and utilization of more complicated non-local analysis techniques in medical imaging and analysis.

4 Future Work

In the future we plan to move away from the concept of separators to clusters. In some sense, separating planes in the data space create areas that can be defined as clusters, but these clusters are restricted to be space-filling and non-overlapping. These restrictions are not required, and performance could improve with their removal.

This work used only two separators. Increasing the number of separators, or clusters, should improve the performance. Using multiple separators provides a way to probe the data space for regions of interest, either because they are important areas for finding cancer or for determining that there is no cancer. The initial analysis has discovered a small region of interest for diagnosing a mammogram as non-cancerous.

This algorithm is not expected to work well on every case because there are too many types of breast cancer and breast characteristics. Tuning the algorithm to the characteristics of the breasts should improve the results. Additionally, letting the algorithm decide when it can do well and when it cannot should also improve performance.

One of the parameters that are learned is the optimal number of suspicious points to use in the analysis. Initial results are always at or near the top of the range that we used, varying from 28 to 31 points depending on the model and weightings. This was surprising because the cancer was usually in the top sixteen if not the top eight points. However, the suspicious points do tend to cluster around a cancer, so including more suspicious points may create a greater distortion of the underlying distribution than fewer points. The learning algorithm does not get the number of points directly, only the cluster differences, so the inclusion of more data should not result in overfitting.

A further improvement might be possible by first transforming the data before filtering, such as applying wavelet analysis to the images before simply thresholding or applying the filter. This has been successfully attempted previously [4] with good results. However, an optimal solution would first combine all of the various filtering and transform methods which create meaningful suspicious points, and then learn an effective analysis from them. This is similar to the effective combination of weak classifiers into a single strong classifier through ensemble learning methods like boosting, which has been successfully used before in tumor classification [24].

5 Conclusion

The overall results of using our techniques are good, our experiments yielded 84% accuracy suggesting that asymmetry is an important measure to incorporate into prescreening or CAD software. Incorporating the technique into CAD shows up to a 71% improvement in the accuracy at a set level of false positives. The technique can be tuned to be more effective at diagnosing cancerous cases, reaching 97% accuracy. We suggest several ways to improve on the methods that we used to measure asymmetry. One method is to convert a mammogram into a connected graph structure of suspicious points and utilize known graph comparison methods for the measure. Another is to use registration of suspicious points from one breast to the other and reduce the suspiciousness of points that have a similar counterpart. Alternatively, increasing the number of separators could improve the method.

Our work has demonstrated the potential of utilizing techniques like image comparisons and other non-local methods with medical imaging. We have shown that we can effectively measure doctor-defined quantities like asymmetry. We believe that in the future, the combination of capturing doctor-defined quantities like asymmetry and machine learning of parameters could be a powerful method for improving the quality of research in medical imaging, and this is one of the avenues of research that we intend to pursue.

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