

# Assignment 4 - Analyzing Functional Ultrasound Images using Tensor Decompositions

## Prerequisites

No particular background knowledge outside the tensor course is required, apart from knowing what correlation is. An interest in biomedical applications might help.

## Background

Functional ultrasound (fUS) is a recently developed neuroimaging technique that measures fluctuations in local blood dynamics induced by changes in neuronal activity [1]. fUS estimates brain activity through the *neurovascular coupling*: this phenomenon dictates that local neuronal activations bring about changes in blood volume in the vessels of the same area. Our focus in this assignment will be capturing responses to sensory stimuli based on the fUS signal using tensor decompositions. fUS typically images a single slice of the brain over time, therefore the data is typically stored as a 3D array  $\underline{\mathbf{X}} \in \mathbb{R}^{N_z \times N_x \times N_t}$ , where the first two modes represent the width and depth of the power Doppler images (PDIs), respectively, and the last mode stands for the time samples of the experiment.

The fUS dataset that we will be working on was acquired from a subject mouse, which was presented with a series of visual stimuli (high-contrast images) during the experimental recording. A specific brain slice was imaged along time, which contains a brain region called the lateral geniculate nucleus (LGN); this region plays a significant role in the processing of visual information. As such, the response within this region is expected to be coupled to the timings of the visual stimulus shown to the subject.

In this assignment, we will first validate this hypothesis through a correlation image. To obtain the correlation image, the Pearson correlation coefficient (PCC) between the (possibly delayed) stimulus time series and the fUS time series of each pixel is computed. The correlation image shows the corresponding PCC for each pixel and is of size  $N_z \times N_x$ . Correlation images provide very important insights while analyzing neuroimaging data, as they demonstrate which regions are activated by the stimulus.

Despite their usefulness, correlation images require that the stimulus is known in advance, which is not always the case: the stimulus time series might not be recorded in the dataset or the experiment might be conducted in the absence of a stimulus (resting state experiment). As an alternative, data-driven techniques such as independent component analysis and tensor decompositions can be used to reveal active brain regions by extracting components that are spatio-temporally coherent, without any a priori information about the stimulus.

In this assignment, you will use the canonical polyadic decomposition (CPD) and the block term decomposition (BTD) to discover task-related components within the brain, i.e., regions that were activated by the stimulus. Two of these regions are the LGN and the visual cortex, but you will also notice some large blood vessels reacting to the stimulus in the correlation image.

To summarize, in this assignment you will first create the correlation image, which can be considered as a guide that shows the regions that we will be searching for using the tensor decompositions. Next, you are asked to compare the performance of the CPD and the BTD when analyzing fUS

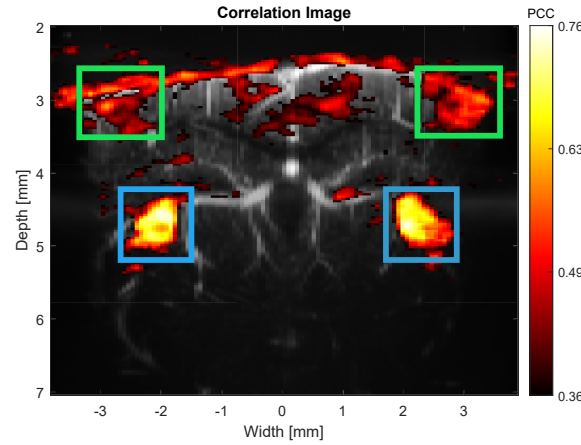
data. Note that as part of this assignment you are expected to implement a BTM algorithm yourself in Matlab. The two files you are asked to fill in are `assignment_4_fUS.m` and `btd_ll1_als_3d.m`.

## Assignment

### Part 1 - Correlation image

The stimulus is represented as a binary signal showing the on-times (value 1) and off-times (value 0) of the visual paradigm (this signal is given to you). To obtain the correlation image, you should calculate the PCC between the pixel time-series and the stimulus signal and display it on an  $N_z \times N_x$  image. Note that the brain response to a given stimulus is often delayed compared to the stimulus onset. As such, the best (i.e., most informative) correlation image is obtained by delaying the stimulus.

In Fig. 1, the correlation image you should come up with is provided for you as a reference. Note that this correlation image only displays the regions that are significantly correlated with the stimulus (with a PCC value above  $\sim 0.3$ ), which are then overlaid on the average PDI (average along the time mode). When you apply a tensor decomposition, you should search for the component(s) whose spatial map(s) resemble the active areas portrayed in the correlation image. These areas include the LGN, the visual cortex, and the large blood vessels on top.

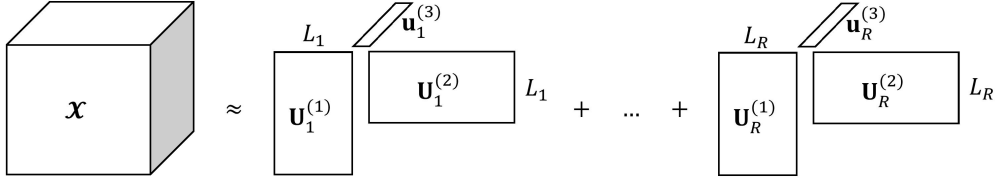


**Figure 1:** Correlation image. The blue and green colored windows show the LGN and visual cortex, respectively. In addition to these regions, the blood vessels on top of the image (in-between the left and right visual cortex) share a significant correlation with the stimulus.

1. To find the optimal delay for plotting the correlation image, you should apply a lag on the stimulus at various values (from 0 to 10 seconds), and calculate the average absolute correlation value over the whole image for each lag. Then, you should determine the lag that maximizes this average, and display the correlation image at this lag. The code that can plot the correlation image has been provided to you (`display_brain_img.m`). You are free to use another method to determine the best delay, as long as you justify your choice.

### Part 2 - Block term decomposition

The BTM can be considered as a generalization of the CPD where the extracted components can have higher multilinear ranks in each mode (CPD components are of rank 1 in all modes). In this assignment, we will be focusing on a particular case of the BTM, known as the  $(L_r, L_r, 1)$ -BTM, which decomposes a 3D input tensor into multilinear rank- $(L_r, L_r, 1)$  terms (Fig. 2). The  $(L_r, L_r, 1)$ -BTM achieves a more general low-rank structure compared to the CPD while preserving uniqueness under relatively mild conditions.



**Figure 2:** Decomposition into multilinear rank- $(L_r, L_r, 1)$  terms.

The  $(L_r, L_r, 1)$ -BTD can be expressed as

$$\underline{\mathbf{X}} = \sum_{r=1}^R \left( \mathbf{U}_r^{(1)} \mathbf{U}_r^{(2)T} \right) \circ \mathbf{u}_r^{(3)}, \quad (1)$$

where  $\circ$  indicates the outer product operation and the matrices  $\mathbf{U}_r^{(1)}$  and  $\mathbf{U}_r^{(2)}$  are of full column rank  $L_r$ . Note that the above equation can also be viewed as a CPD by rewriting it as follows

$$\underline{\mathbf{X}} = \sum_{r=1}^R \sum_{l=1}^{L_r} \mathbf{U}_r^{(1)}(:, l) \circ \mathbf{U}_r^{(2)}(:, l) \circ \mathbf{u}_r^{(3)}. \quad (2)$$

Thus, similar to the CPD, the  $(L_r, L_r, 1)$ -BTD can be solved for using an alternating least squares (ALS) approach.

1. Fill in the code that implements the ALS algorithm for the  $(L_r, L_r, 1)$ -BTD in `btd_lll_als_3d.m`.  
*Hint:* You can look in the literature for guidance in this task.

### Part 3 - Problem formulation

1. Give a short problem definition that describes the problem we are solving and the methods we are investigating (no math should be included). You are encouraged to include specific research questions. Do not copy the assignment text for this.

### Part 4 - Applying and comparing the CPD and the $(L_r, L_r, 1)$ -BTD

Apply the CPD on the fUS data (you can use the assignment 2 solution file `cpd_als_3d.p` provided to you) and answer the following questions:

1. Are you able to extract a component whose spatial map points to any of the expected areas?  
*Hint 1:* You should combine the factor matrices of the first two modes to construct a spatial map. Do not forget to use a colorbar!  
*Hint 2:* Keep in mind the sign ambiguity of the CPD.
2. If your answer to the previous question is yes, what information does the temporal signature of that component entail? Is the temporal signature significantly correlated to the stimulus (i.e., with a PCC value above 0.3)?
3. How did you determine the number of sources? Justify your choice.
  - Are you still able to extract a meaningful component when you select it differently?
  - What happens when you run the CPD with different initializations and a fixed number of sources?

Now apply the  $(L_r, L_r, 1)$ -BTD and answer the following questions:

1. The first and second question as in the CPD case above.
2. While applying the BTM, which mode of the fUS tensor did you select to be rank-1 and why?
3. Compared to the CPD, what kind of differences do you observe in the extracted component(s) of interest?
4. What do you think is the reason for your observations in the previous question?  
*Hint:* Think of the assumptions made by the CPD and the BTM.
5. How did you determine the number of sources and the factor matrices ranks  $L_r$ ?
  - Are you still able to extract a meaningful component when you select them differently?
  - What happens when you run the BTM with different initializations and a fixed number of sources?

## Part 5 - Conclusions

Please describe your final conclusions given your observations in the previous parts.

## Tips

- You can check [2] for inspiration for answering the questions above.
- Some functions from previous assignments that could be useful have been provided to you. You are free to include any other function you code, as long as you explain its use.

## Consultation meetings

1. In meeting no. 1, you are expected to have completed parts 1-3 of the assignment, with special emphasis on parts 1 and 3. If part 2 is not completed, you are expected to at least have identified the relevant literature and started coding the ALS algorithm.
2. Meeting no. 2 is reserved for any final questions or clarifications.

## Grading rubric

You will be graded based on how well you follow the guidelines of the assignment, as well as your observations and critical thinking. In particular, the distribution of the 20 points is as follows:

- Correlation image [1 point],
- $(L_r, L_r, 1)$ -BTM implementation [5 points],
- Problem formulation [2 points],
- Applying the CPD and answering the corresponding questions [6 points],
- Applying the BTM and answering the corresponding questions [6 points],
- Bonus points can be earned by means of additional discussions or experiments.

Your report should follow the attached IEEE conference template and not exceed 6 pages. Ensure all your plots are visible and clear (do not include the images as .png files but as .eps or .pdf files!!).

## References

- [1] E. Macé, G. Montaldo, I. Cohen, M. Baulac, M. Fink, and M. Tanter, “Functional ultrasound imaging of the brain,” *Nature Methods*, vol. 8, pp. 662–664, 2011.
- [2] C. Chatzichristos, E. Kofidis, M. Morante, and S. Theodoridis, “Blind fMRI source unmixing via higher-order tensor decompositions,” *Journal of Neuroscience Methods*, vol. 315, pp. 17–47, 2019.