

## Functional Magnetic Resonance in Binasal Hemianopia

Jan Lestak<sup>1,2,3\*</sup>, Jaroslav Tintera<sup>1</sup>, Jiri Zahlava<sup>1</sup>, Martin Sverepa<sup>1</sup> and Pavel Rozsival<sup>3</sup>

<sup>1</sup>JL Clinic, V Hurkach 1296/10, Prague, Czech Republic

<sup>2</sup>Czech Technical University in Prague (Faculty of Biomedical Engineering), Czech Republic

<sup>3</sup>Charles University (Faculty of Medicine in Hradec Kralove), Czech Republic

\*Corresponding author: Jan Lestak, JL Clinic, V Hurkach 1296/10, 158 00 Prague 5, Czech Republic, E-mail: lestak@seznam.cz

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### Abstract

The authors report a case of binasal hemianopia in a 34-year-old female who, at the age of 22 years, underwent resection of a pinealoma located at the 3rd ventricle. Less than a year after the surgery, partial visual field losses occurred. Examination of retina, retinal nerve fiber layer (RNFL) as well as examination of the ganglion cell complex (GCC) excluded a pregeniculate lesion. Functional magnetic resonance imaging (fMRI) responses evoked by both bi- and monocular stimulation confirmed our finding of reduced activity in the visual cortex corresponding to the damage in a portion of the visual pathway.

**Keywords:** Complete binasal hemianopia; fMRI; RNFL; GCC; Pineal region tumor; Third cerebral ventricle

### Introduction

The existence of complete binasal hemianopia has been disputed since the introduction of the first perimeters. In 1877, Foerster, one of its inventors, wrote about the relationship between the nervous system and the visual organ in the first edition of the Graef-Saemisch

Compendium: "there is no binasal hemianopia" [1]. After the introduction of automated static perimetry, the results of visual field changes became more precise and a large number of incomplete binasal hemianopia's were eliminated. In 2011, we reported on a patient with complete binasal hemianopia [2]. Due to the fact that this report has been questioned, we decided to reopen the case, provide additional photo documentation and verify the visual field changes using functional magnetic resonance (fMRI). We want to prove that the change in activation laterality when comparing the visual stimulation of the right and left eye separately can be detected by fMRI and support results of the standard examination of the patient with binasal hemianopia. Therefore, we formulate the following hypothesis: A patient with binasal hemianopia has weaker activation in the right occipital lobe during stimulation of the right eye and weaker activation in the left occipital lobe during stimulation of the left eye. Additionally, the patient can have weaker activation generally in both hemispheres compared to healthy controls. This fact leads to evaluate lateralization index (LI) of the activation rather than to compare only strength of the activation expressed in number of statistically significant voxels [3].

### Material and methods

#### Patient history

In November 2002, a 21-year-old nurse suddenly lost consciousness during her night shift, accompanied by limb cramps, without wetting or biting. A similar situation repeated itself in December 2002. During her examination in December 2002, she was conscious, cooperative,

well-oriented, with a slight bilateral ataxia and medium-grade paleocerebellar symptomatology. The ocular finding was bilaterally normal including the perimeter.

The MRI examination of the brain carried out on December 16, 2002 revealed a hypoinvasive structure in the pineal region, sized 21 × 17 × 19 mm, slightly ventrocaudally compressing the quadrigeminal bodies, hyperintensive in the T2 weighing, with a narrow non-enhancing border; the ventricular system was narrow and the central line was without any overpressure. The pituitary stalk was localized behind the chiasm. On January 14, 2003 a tumor was removed from the pineal region through suboccipital craniotomy. The tumor was gradually sharply separated from the base, respecting its boundary against the upper stalk, and was radically removed. The histological examination established a pineocytoma.

The postoperative MRI of the brain (January 16, 2003) confirmed the radical removal of a pinealoma with no obstruction of the liquor pathways. Eighteen days after surgery, the patient was released into home care. She did not appear to have any problems, apart from slight paleocerebellar symptomatology [2].

#### Outpatient follow-up

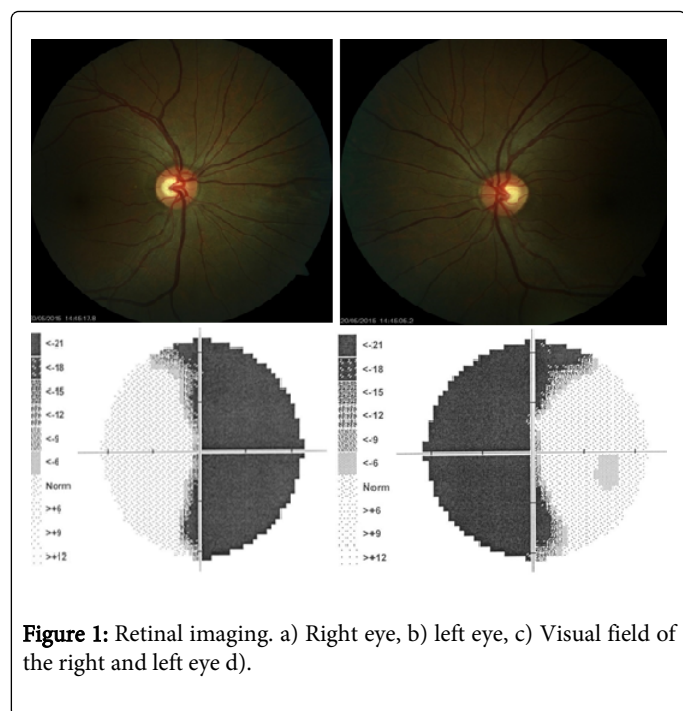
The patient's neurological state did not change until December 2003, including normal visus and provisionally examined perimeter. The ocular findings at the follow-up examination carried out by an ophthalmologist in April 2003 were normal, including the objectively performed perimeter examination. On May 27, 2003 the patient suffered a consciousness disorder (approx. 12 minutes), with clonic-tonic spasms and wandering motion of the eyes. During her hospitalization in the Neurological Department, her problem did not recur. ECG, examination of the eyegrounds and laboratory examination showed no pathological findings. CT and MRI examinations proved reparative gliosis around the aqueduct, otherwise without any other pathology. Early in December 2003, the patient experienced subjectively impaired visus. The eye examination revealed increased intraocular pressure. During her hospitalization in the Eye

Clinic, the first scotomas in the nasal halves of both visual fields were found on the perimeter, further to the right.

The monthly follow-up examinations of the perimeter showed that, until the end of June 2004, the finding progressed gradually into complete binasal hemianopia, and has persisted until now. In July 2004, the patient was examined at the lead author's clinic, where the examination of electrical retinal functions did not reveal any changes in the flash or pattern electroretinograms. The next ophthalmological examination was carried out in November 2010. Both the external and intraocular findings were normal, while the binasal hemianopia in the perimeter still persisted. Repeated MRI examinations of the brain (annually until 2010) did not show any recurrence of the pinealoma, the reactive changes disappeared (=reactive gliosis) and the cerebral finding was normal.

The last ophthalmological examination was performed in May 2015, with a bilaterally normal ocular finding: VA with -3.5, -2.5 respectively=-1.0; excavation of the fundus with C/D ratio of 0.4; IOPs were 18 and 18 mmHg; perimeter: complete binasal hemianopia.

This finding has remained unchanged since December 2003. Figure 1 shows normal retina. Figure 2 shows nerve fiber layer and ganglion cell complex.



**Figure 1:** Retinal imaging. a) Right eye, b) left eye, c) Visual field of the right and left eye d).

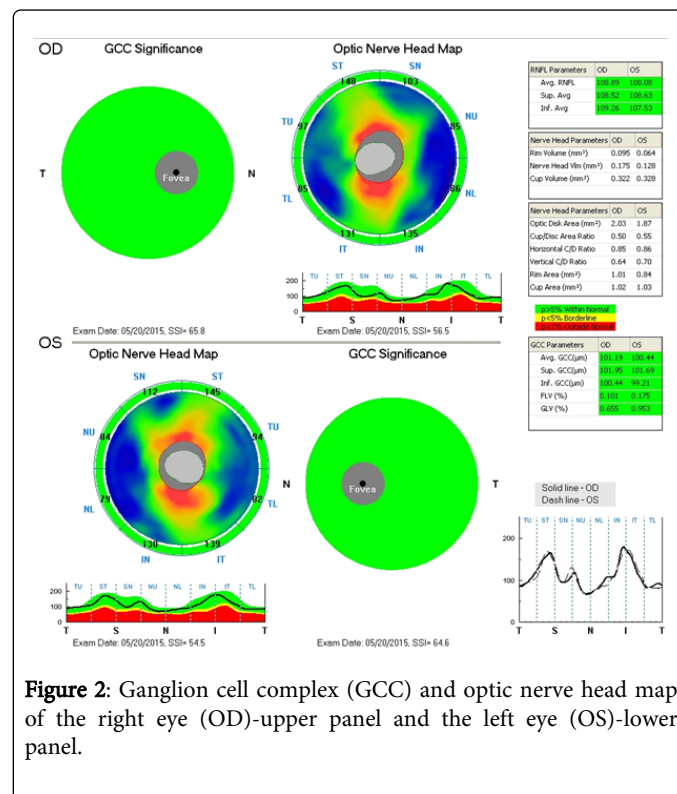
### Control group

Control group consists of 6 healthy female subjects (average age of 35 years). All subjects had complete ophthalmologic examination excluding any pathology in visual field and attended the fMRI examination with identical protocol as the patient.

### Functional MR imaging (fMRI)

Functional MRI examinations were carried out on the Philips Achieva 3T TX MR system (Philips Healthcare, Eindhoven, Netherlands) with a magnetic field strength of 3 Tesla, using the blood oxygen level dependent (BOLD) contrast. A standard 32-channel

SENSE head RF coil was used for scanning. For fMRI measurements based on the BOLD technique, the gradient-echo EPI sequence was used with the following parameters: TE=30 ms, TR=3 s, flip angle of 90°. The measured volume contained 39 continuous slices. The voxel size measured was 2 × 2 × 2 mm (FO=208 × 208 mm, matrix 104 × 104, SENSE factor 1.8).



**Figure 2:** Ganglion cell complex (GCC) and optic nerve head map of the right eye (OD)-upper panel and the left eye (OS)-lower panel.

Optical stimulation was performed by a black/white checkerboard alternated with its negative image with a frequency of 2 Hz. The visual size of the black and white checkerboard was 25.8 × 16.2 degrees. The measurements consisted of a sequence of five 30-second active phase periods and five resting periods of the same length (each of 10 dynamic scans). During the resting phase, a static crosshair situated in the center of the visible field was projected for the view fixation. In total, every measurement included 100 dynamic scans and took 5 minutes. Each eye was examined by means of separate fMRI measurement (LE, RE) and also one control measurement was performed by stimulating both eyes together (LE+RE) (Figure 3).

The obtained data were processed using SPM8 software and general linear model (GLM). During the pre-process, the data were motion corrected (realignment), corrected for time-shift of individual slices (slice timing), then smoothed using a Gaussian filter with FWHM of 6 × 6 × 6 mm and finally normalized into the MNI\_152 space. For statistical evaluation, the GLM with canonical hemodynamic response function (HRF) applied to the block scheme of the stimulation was used. Statistical maps were thresholded at the level of p=0.05 with FWE correction for multiple observations. The extent of the activation (number of statistically significant voxels) is strongly dependent on the choice of the statistical threshold. In case of very different strength of the activation the evaluation of the activation laterality can be challenging. This is especially valid when comparing a patient (with general weaker activation) with healthy controls. Therefore, a threshold independent method to evaluate lateralization of the fMRI

activation using bootstrap introduced by Wilke et al. was used and the extent of activations in occipital lobe of both hemispheres is expressed as lateralization index (LI) which is a ration defined by the relation.

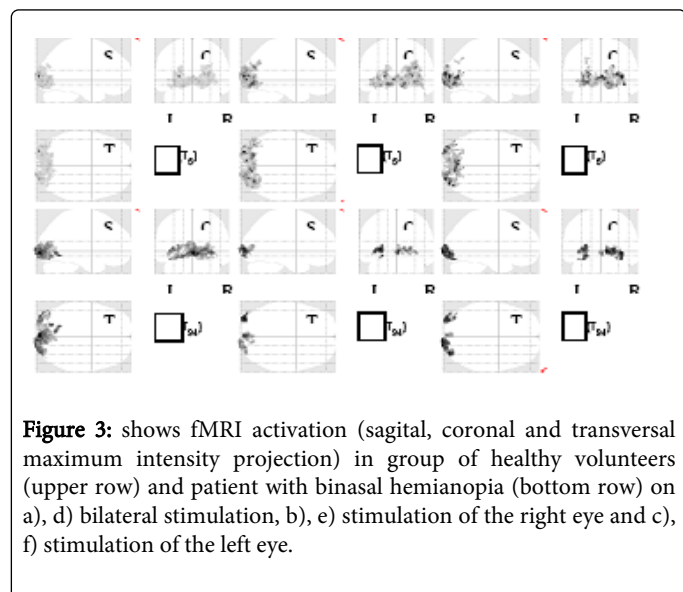
$$LI = (ALH - ARH) / (ALH + ARH)$$

Where ALH, ARH is the number of activated voxels in the left or right hemisphere resp. Complete left side activation is represented by LI=1 while LI=-1 means exclusively right side activation.

Lateralization index was calculated by bootstrap method in SPM toolbox [4,5]. Default parameters of the LI toolbox were used for the calculation: sub-sample size k=25%, minimum sample size 5, maximum sample size 10 000 and only voxels from occipital lobes were used by masking.

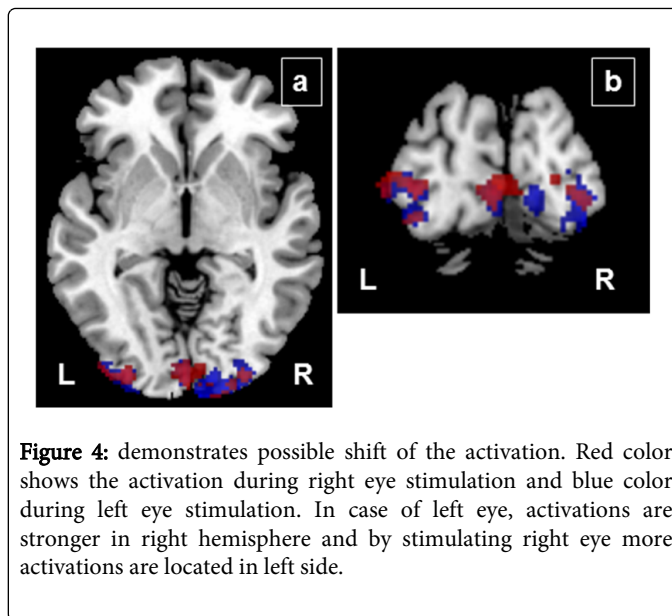
## Results

The comparison of activations between the group of 6 healthy female volunteers (a-c) and patient with binasal hemianopia (d-e) is shown in (Figure 3). In average, healthy subjects had more extensively activated visual cortex than the patient. Moreover, difference in the laterality of the activation can be seen also in (Figure 4) where red color represents the response on the stimulation of the right eye and blue color the activation during left eye stimulation. Decrease of the activation in the right occipital lobe as a response to right eye stimulation as well as a decrease in the left occipital lobe during left eye stimulation is visible.



**Figure 3:** shows fMRI activation (sagittal, coronal and transversal maximum intensity projection) in group of healthy volunteers (upper row) and patient with binasal hemianopia (bottom row) on a), d) bilateral stimulation, b), e) stimulation of the right eye and c), f) stimulation of the left eye.

More quantitative approach using the quantification of LI is shown in Table 1. Results of LI during bilateral, right and left eye stimulation are presented for all 6 healthy controls and for the patient with binasal hemianopsia. All volunteers have either left side activation dominance (LI is positive) or right side dominance (negative LI) for both eyes but the patient has positive LI=0.18 for right eye stimulation (decreased right hemisphere activation) and negative LI=- 0.16 for left eye stimulation (decreased left hemisphere activation).



**Figure 4:** demonstrates possible shift of the activation. Red color shows the activation during right eye stimulation and blue color during left eye stimulation. In case of left eye, activations are stronger in right hemisphere and by stimulating right eye more activations are located in left side.

Subject	LI bilateral stimulation	LI right stimulation eye	LI left stimulation eye
control 1	0.32	0.29	0.41
control 2	-0.21	-0.47	-0.08
control 3	-0.29	-0.26	-0.14
control 4	0.30	0.25	0.38
control 5	0.31	0.30	0.20
control 6	-0.15	-0.01	-0.09
patient BH	0.16	0.18	-0.16

**Table 1:** shows the activation laterality expressed as a lateralization index (LI) for 6 healthy female volunteers and patient with binasal hemianopia during bilateral stimulation and separate stimulation of the right or left eye.

## Discussion

Binasal hemianopia is a condition structurally related to uncrossed optic fibers. This means that, during stimulation of either of the eyes, the signal alteration should occur in the ipsilateral occipital hemisphere.

Due to the extent of the damage involving not only nasal but partly also temporal visual fields, the activity of the visual cortex during bi- as well as monocular stimulation in our patient is markedly lower.

The exact localization and development process of the lesion remain elusive. If the binasal hemianopia had a structural correlate in the peripheral portion of the visual pathway (pregeniculate), pathology would be found at the optic discs, in the nerve fiber layer of the retinal ganglion cells, as well as in the ganglion cells themselves. In our patient, no such pathologies were observed. It is feasible, however, to assume a central (suprageniculat) localization of the lesion.

The decreased activity of the visual cortex supports the presence of a structural, rather than a functional correlate of the binasal changes.

## Conclusion

By fMRI, we were able to confirm a structural background for the binasal hemianopia in our patient, although we can only speculate on the detailed localization of the lesion.

The study protocol was approved by the local Ethics Committee and the study was performed in accordance with Good Clinical Practice and the Declaration of Helsinki.

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## Conflict of interest statement

The authors state that there are no conflicts of interest regarding the publication of this article.

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