

UNIWERSYTET TECHNOLOGICZNO-PRZYRODNICZY
IM. JANA I JĘDRZEJA ŚNIADECKICH W BYDGOSZCZY
ZESZYTY NAUKOWE NR 260
TELEKOMUNIKACJA I ELEKTRONIKA 15 (2011), 49-61

MRI IMAGE ANALYSIS IN PATIENTS WITH A TUMOR OF THE CENTRAL NERVOUS SYSTEM – AN ATTEMPT OF DEVELOPING A MANAGEMENT ALGORITHM

Maciej Śniegocki¹, Agnieszka Nowacka², Jacek Fisz³
Aleksandra Śniegocka⁴, Marcin Buczkowski⁵

¹ Department of Neurotraumatology CM UMK

² Department of Neurosurgery and Neurotraumatology CM UMK

³ Department of Computer Science and Methodology of Science Work CM UMK

⁴ Department of Clinical Psychology CM UMK
University Hospital No. 1, ul. M. Skłodowskiej-Curie 9, 85-094 Bydgoszcz

⁵ Department of Physics and Biophysics UWM,
Plac Cieszyński 1 (bl. 43), 10-726 Olsztyn

Summary: Magnetic resonance imaging study is currently the reference method for the detection and diagnosis of the central nervous system tumors. A large number of tumors, especially high-grade, has a higher water content in the cells, which results in prolongation of MRI T1 and T2 what appearance as increased signal intensity in in T2-weighted images and the reduction in T1-weighted images. MRI can be performed with administration of contrast agent, which shortens T1 and increases signal on T1-weighted sequences. This allows to identify areas of increased angiogenesis), which is the exponent of the cancer malignancy degree and its biological activity. Obtained MRI images are analyzed and evaluated by a radiologist and a clinician. Most of the time it is the "by the eye" analysis, which is based on the MRI image evaluation by the generally accepted radiological standards. However, this method is relatively inaccurate. which in turn can bring to the wrong diagnosis of the disease and implementation or even lack of implementation of appropriate treatment. More and more researches are conducted in this area, but developed methods are usually very complicated and difficult to carry out by the "layman" which is the clinician. That is why the attempt is made, to develop a simple and clear algorithm for MRI image analysis in patients with the central nervous system tumors, allowing for quick and objective evaluation of magnetic resonance imaging study.

Keywords: tumor, MRI image, angiogenesis, image analysis

1. TUMORS OF THE CENTRAL NERVOUS SYSTEM

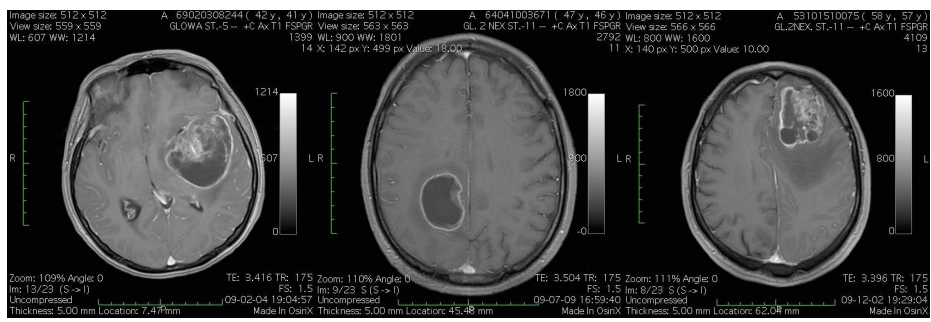
Central nervous system tumors are approximately 1.5% of all cancers occurring in the human body, but their incidence is still increasing. Globally, the incidence of deaths due to intracranial tumors is estimated to be approximately 2.3% of all deaths due to cancer. 70-75% of them are primary tumors, and 20-25% are metastatic tumors. Due to the specific structure and functions of the brain, central nervous system tumors differ

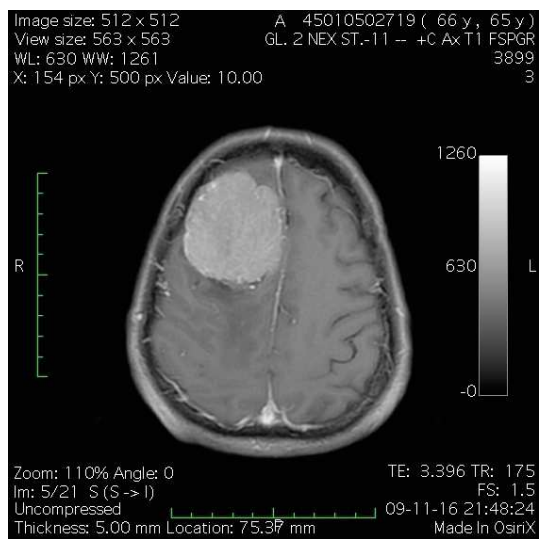
significantly from tumors located in other parts of the body. The very location of the tumor, regardless of its growth and malignancy, affects on the prognosis and patient's functional status. In case of location of the lesion in the area of important vital structures, both a tumor growth and its surgical treatment is a huge risk of severe disability or death of the patient.

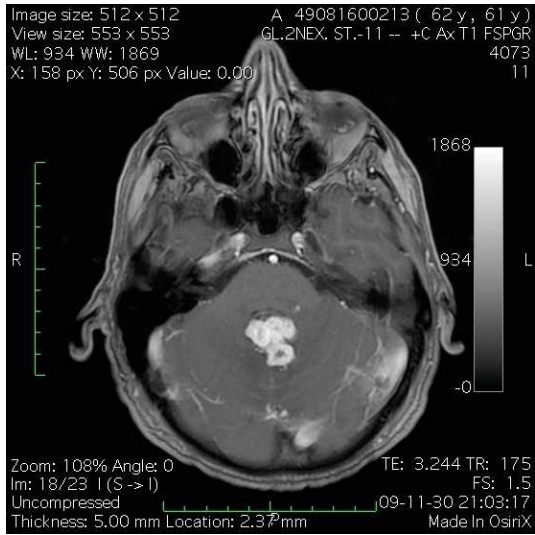
Tumors of the central nervous system can be classified according to several basic criteria. The first one is the presentation of tumors due to their location within the cranial cavity. There are supratentorial tumors (located in the lobes of the brain, in deep structures of the telencephalon and diencephalon), which are 80-85% of all intracranial tumors in adults, and 40% in children, and infratentorial tumors (located within the cerebellum, brainstem, or the cerebellopontine angle), representing 15-20% of intracranial tumors in adults and 60% in children. This division is based on images obtained in radiological examinations, such as conventional radiography (X-ray), computed tomography (CT), magnetic resonance imaging (MRI) or positron emission tomography (PET). It is widely used in clinical practice, mainly because the location of the pathology is responsible for the occurrence of specific neurological symptoms, and allows to determine the quality of possible damage of the central nervous system after the surgery. Another criterion of division of the central nervous system tumors is the origin of the tumor tissue, first introduced by the World Health Organization (WHO) in 1979. The latest, 2007, WHO classification accounts the current state of knowledge about the biology of cancer, intracellular regulatory pathways, disturbed in the process of oncogenesis, and aspects of the therapeutic methods prediction.

TUMORS OF NEUROEPITHELIAL TISSUE	Desmoplastic infantile astrocytoma/ganglioglioma	Malignant peripheral nerve sheath tumour (MPNST)	Angiosarcoma
Astrocytic tumours	Dysembryoplastic neuroepithelial tumour	- Epithelioid MPNST	Kaposi sarcoma
Piloicytic astrocytoma	Gangliocytoma	- MPNST with mesenchymal differentiation	Ewing sarcoma (PNET)
- Piloxyoid astrocytoma	Ganglioglioma	- Melanotic MPNST	Primary melanocytic lesions
Subependymal giant cell astrocytoma	Anaplastic ganglioglioma	- MPNST with glandular differentiation	Diffuse melanocytosis
Pleomorphic xanthoastrocytoma	Central neurocytoma		Melanocytoma
Diffuse astrocytoma	Extraventricular neurocytoma	TUMOURS OF THE MENINGES	Malignant melanoma
- Fibrillary astrocytoma	Cerebellar liponeurocytoma		Meningeal melanomatosis
- Gemistocytic astrocytoma	Papillary glioneuronal tumour [PGNT]	Tumours of meningeothelial cells	Other neoplasms related to the meninges
- Protoplasmic astrocytoma	Rosette-forming glioneuronal tumour of the fourth ventricle	Meningioma	Haemangioblastoma
Anaplastic astrocytoma	Paraganglioma	- Meningothelial	LYMPHOMAS AND HAEMATOPOIETIC NEOPLASMS
Glioblastoma		- Fibrous (fibroblastic)	Malignant lymphomas
- Giant cell glioblastoma	Tumours of the pineal region	- Transitional (mixed)	Plasmocytoma
- Gliosarcoma	Pineocytoma	- Psammomatous	Granulocytic sarcoma
Gliomatosis cerebri	Pineal parenchymal tumour of indeterminate differentiation	- Chordoid	GERM CELL TUMOURS
Oligodendroglial tumours	Pineoblastoma	- Clear cell	Germinoma
Oligodendroglioma	Papillary tumour of the pineal region [PTPR]	- Atypical	Embryonal carcinoma
Anaplastic oligodendroglioma		- Papillary	Yolk sac tumour
Oligoastrocytic tumours	Embryonal tumours	- Rhabdoid	Choriocarcinoma
Oligoastrocytoma	Medulloblastoma	- Anaplastic (malignant)	Teratoma
Anaplastic oligoastrocytoma	- Desmoplastic/nodular medulloblastoma	Mesenchymal tumours	- Mature
Ependymal tumours	- Medulloblastoma with extensive nodularity	Lipoma	- Immature
Subependymoma	- Anaplastic medulloblastoma	Angiolipoma	- Teratoma with malignant transformation
Myxopapillary ependymoma	- Large cell medulloblastoma	Hibernoma	Mixed germ cell tumour
Ependymoma	CNS primitive neuroectodermal tumour	Liposarcoma	TUMOURS OF THE SELLAR REGION
- Cellular	- CNS Neuroblastoma	Solitary fibrous tumour	Craniopharyngioma
- Papillary	- CNS Ganglioneuroblastoma	Fibrosarcoma	- Adamantinomatous
- Clear cell	- Medulloepithelioma	Malignant fibrous histiocytoma	- Papillary
- Tanycytic	- Ependymblastoma	Leiomyoma	Granular cell tumour
Anaplastic ependymoma	Atypical teratoid/rhabdoid tumour	Leiomyosarcoma	Pituitaryoma
Choroid plexus tumours	TUMOURS OF CRANIAL AND PARASPINAL NERVES	Rhabdomyoma	Spindle cell oncocytoma of the adenohypophysis
Choroid plexus papilloma	Schwannoma (neurilemoma, neurinoma)	Rhabdomyosarcoma	METASTATIC TUMOURS
Atypical choroid plexus papilloma	- Cellular	Chondroma	
Choroid plexus carcinoma	- Plexiform	Chondrosarcoma	
Other neuroepithelial tumours	- Melanotic	Osteoma	
Astroblastoma	Neurofibroma	Osteosarcoma	
Choroid glioma of the third ventricle	- Plexiform	Osteochondroma	
Angiocentric glioma	Perineurinoma	Haemangioma	
Neuronal and mixed neuronal-glial tumours	- Perineurinoma, NOS	Epithelioid haemangiopericytoma	
Dysplastic gangliocytoma of cerebellum (Lhermitte-Duclos)	- Malignant perineurinoma	Haemangiopericytoma	
		Anaplastic haemangiopericytoma	

Fig. 1. Classification of the central nervous system tumors according to WHO, 2007.







rearranges relatively to the magnetic field lines. Then they are stimulated by the resonant frequent electromagnetic pulses. During the stimulation decay, occurs an emission of a radio frequency waves. Received signals differ by intensity, depending on the type of tissue from which they originate. A large number of tumors, especially high-grade, has a higher water content in the cells, which results in prolongation of MRI T1 and T2 what appear as increased signal intensity in T2-weighted images and the reduction in T1-weighted images.

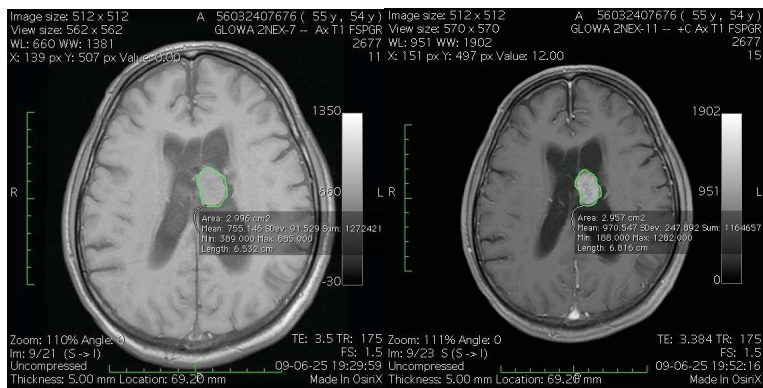
Table 2. Type of MRI signal in the case of selected central nervous system tumors.

Tumor histological type	T1 – weighted images	T2 – weighted images
Low grade gliomas	hipo/-izointensive	hiperintensive
High grade gliomas	hipo/-izointensive	hiperintensive
Meningiomas	iso-/hipointensive	iso-/hipointensive
Metastatic tumors	hipo/-izointensive	hiperintensive

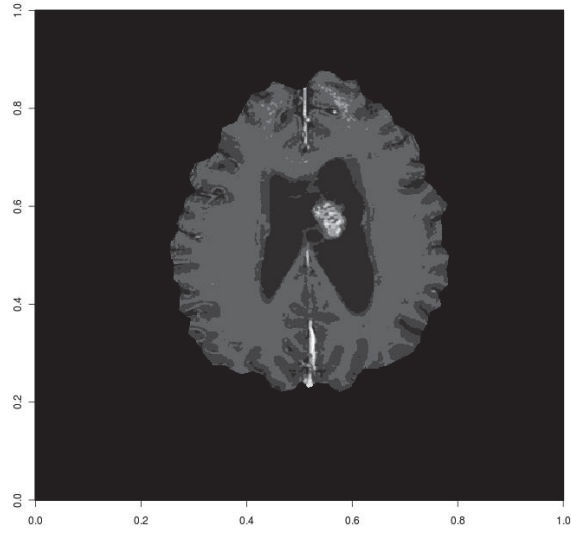
As with CT, MRI can be performed without or with administration of contrast agent (usually gadolinium), which shorten T1 and increase signal on T1-weighted sequences. This allows to identify areas of increased blood flow (increased angiogenesis). Increased angiogenesis, the formation of blood vessels in tumor tissues, is the exponent of the cancer malignancy degree and its biological activity. Microcirculation can be illustrated by examination of brain tissue perfusion (eg. PWI MRI – weighted perfusion MRI, DCE MRI - dynamic contrast enhanced MRI). Administration of the contrast agent can also detect damage to the blood-brain barrier and allows to find the boundary between tumor and surrounding edema. The degree of after-contrast signal increase depends on the concentration of contrast agent and magnetic fields force.

There are many MRI sequences and techniques, eg. FLAIR – fluid attenuated inversion recovery, STIR – short tau inversion recovery, MRA – magnetic resonance angiography, DWI – diffusion weighted imaging, MRS – magnetic resonance spectroscopy, DTT MRI – diffusion tensor tractography MRI. This an examination highly exceed computed tomography in the diagnosis of tumors of the central nervous system. Of course it has its limitations, such as the relatively long examining time, sensitivity to motion artifacts, lack of examination possibility in case of the metal presence in patient's body, and the high price of examination. Despite that, it has many advantages, that put it in the first place in neuroimaging diagnosis, including:

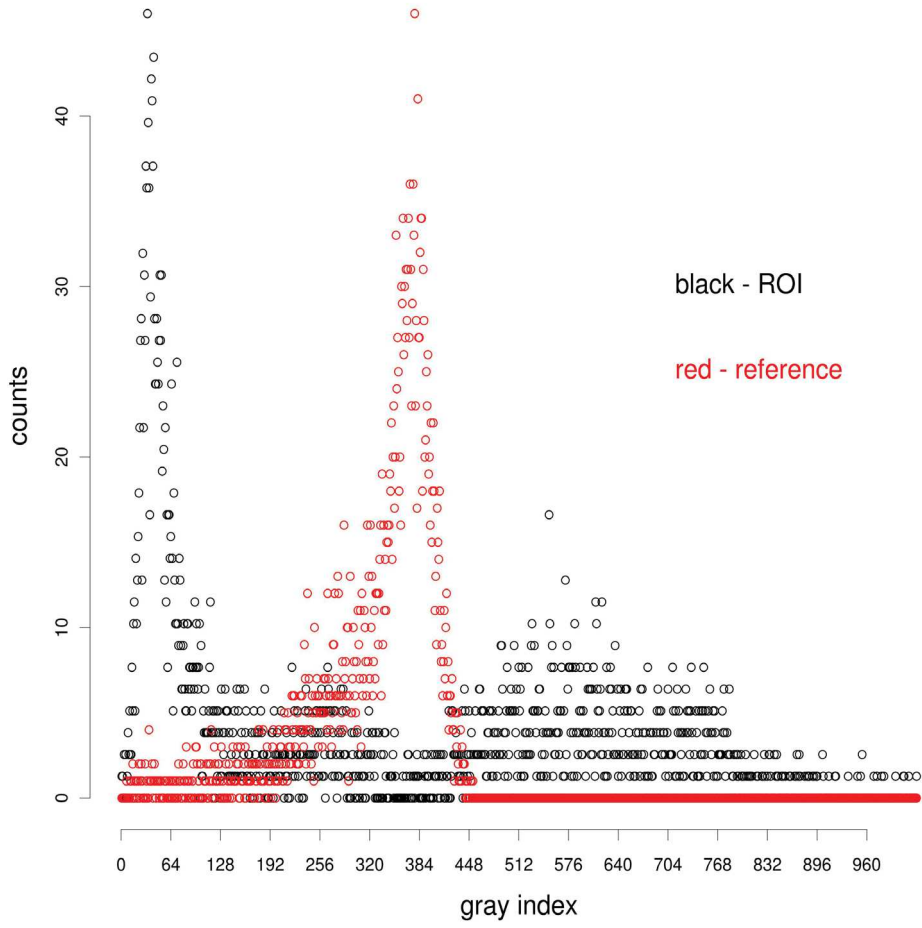
- the possibility of obtaining images in any plane, which allows to define a very precise topographical relationships in relation to adjacent anatomical structures,
- high contrast resolution between imaging tissues,
- lack of bone artifacts,
- greater sensitivity than CT,
- possibility to conduct the examination in pregnant women (very low risk of adverse effects on the fetus of the magnetic field, as opposed to harmful ionizing radiation X in the case of CT),
- non-invasive examination,
- repeatability of examination.



clustered dpm classification



1024 gray levels



The new method can help to facilitate the work of both radiologists – the assessment of MRI images, and clinicians – when planning treatment, monitoring the disease and predicting prognosis for patient survival. Due to its simplicity and fast, direct obtaining of concrete results, it can also be used by those less experienced in the analysis of magnetic resonance images.

BIBLIOGRAPHY

- [1] Aksoy F.G., Lev M.H., 2000. Dynamic Contrast – Enhanced Brain Perfusion Imaging: Technique and Clinical Applications. *Sem Ultrasound CT MRI*, 21 (6), pp. 462-477.
- [2] Al-Okaili R.N., Krejza J., Woo J.H., Wolf R.L., O'Rourke D.M., Judy K.D., Poptani H., Melhem E.R., 2007. Intraaxial brain masses: MR imaging-based diagnostic strategy – initial experience. *Radiology* 243, pp. 539-550.
- [3] Aronen H.J., Gazit I.E., Louis D.N., Buchbinder B.R., Pardo F.S., Weisskoff R.M., Griffith R.H., Cosgrove G.R., Halpern E.F., Hochberg F.H., Rosen B.R., 1994. Cerebral Blood Volume Maps of Gliomas: Comparison with Tumor Grade and Histologic Findings. *Radiology* 191, pp. 41-51.
- [4] Brem S., Cotran R., Folkman J., 1972. Tumor angiogenesis: a quantitative method for histologic grading. *J Natl Cancer Inst* 48, pp. 347-356.
- [5] Emblem K.E., Zoellner F.G., Tennoe B., Nedregaard B., Nome T., Due-Tonnessen P., Hald J.K., Scheie D., Bjornerud A., 2008. Predictive modeling in glioma grading from MR perfusion images using support vector machines. *Magn Reson Med* 60, pp. 945-952.
- [6] Folkman J., 1992. The role of angiogenesis in tumor growth. *Semin Cancer Biol*, 3, pp. 65-71.
- [7] Folkman J., 1990. What is the evidence that tumors are angiogenesis dependent? *J Natl Cancer Inst* 82, pp. 4-6.
- [8] Ginsberg L., Fuller G., Schomer D., Kau B.A., Kispert D.B., 1996. Does lack of enhancement of brain tumors on MR imaging correlate with low grade malignancy? A histopathologic study. [In:] *Proceedings of the American Society of Neuroradiology, Seattle-Washington*, pp. 32-33.
- [9] Glotsos D., Tohka J., Ravazoula P., Cavouras D., Nikiforidis G., 2005. Automated diagnosis of brain tumors astrocytomas using probabilistic neural network clustering and support vector machines. *Int J Neural Syst* 15, pp. 1-11.
- [10] Greenberg M.S., Arredondo N., 2010. *Handbook of neurosurgery – 7th edition*, Thieme.
- [11] Higano S., Yun X., Kumabe T., Watanabe M., Mugikura S., Umetsu A., Sato A., Yamada T., Takahashi S., 2006. Malignant astrocytic tumors: clinical importance of apparent diffusion coefficient in prediction of grade and prognosis. *Radiology* 241, pp. 839-846.
- [12] Huang Y., Lisboa P.J.G., El-Deredy W., 2003. Tumour grading from magnetic resonance spectroscopy: a comparison of feature extraction with variable selection. *Stat Med* 22, pp. 147-164.
- [13] Knopp E.A., Cha S., Johnson G., Mazumdar A., Golfinos J.G., Zagzag D., Miller D.C., Kelly P.J., Kricheff I.I., 1999. Glial Neoplasms: Dynamic Contrast – Enhanced T2* – weighted MR Imaging. *Radiology* 211, pp. 791-798.

- [14] Kremer S., Grand S., Remy C., Esteve F., Lefournier V., Pasquier B., Hoffmann D., Benabid A.L., Le Bas J.F., 2002. Cerebral blood volume mapping by MR imaging in the initial evaluation of brain tumors. *J Neuroradiol* 29, pp. 105-113.
- [15] Law M., Cha S., Knopp E. A., Johnson G., Arnett J., Litt A.W., 2002. High-Grade Gliomas and Solitary metastases: Differentiation by Using Perfusion and Proton Spectroscopic MR Imaging. *Radiology* 222, pp. 715-721.
- [16] Lev M.H., Hochberg F., 1998. Perfusion Magnetic Resonance Imaging to Assess Brain Tumor Responses to New Therapies. *Cancer Control* 5 (2), pp. 115-123.
- [17] Lev M.H., Ozsunar Y., Henson J.W., Rasheed A.A., Barest G.D., Harsh G.R., Fitzek M.M., Chiocca E.A., Rabinov J.D., Csavoy A.N., Rosen B.R., Hochberg F.H., Schaefer P.W., Gonzalez R.G., 2004. Glial tumor grading and outcome prediction using dynamic spin-echo MR susceptibility mapping compared with conventional contrast-enhanced MR: confounding effect of elevated rCBV of oligodendrogliomas. *Am J Neuroradiol* 25, pp. 214-221.
- [18] Liberski P.P., Kozubski W., Biernat W., Kordek R., 2011. *Neuroonkologia kliniczna*. Wyd. Czelej.
- [19] Louis D.N., Ohgaki H., Wiestler O.D., Cavenee W.K., Burger P.C., Jouvet A., Scheithauer B.W., 2007. *The 2007 WHO Classification of Tumours of the Central Nervous System*. Springer-Verlag.
- [20] Lüdemann L., Grieger W., Wurm R., Budzisch M., Hamm B., Zimmer C., 2001. Comparison of dynamic contrast-enhanced MRI with WHO tumor grading for gliomas. *Eur Radiol* 11, pp. 1231-1241.
- [21] Majó's C., Julia-Sape M., Alonso J., Serrallonga M., Aguilera C., Acebes J.J., Arús C., Gili J., 2004. Brain tumor classification by proton MR spectroscopy: comparison of diagnostic accuracy at short and long TE. *Am J Neuro-radiol* 25, pp. 1696-1704.
- [22] Melhem E.R., Davatzikos C., 2008. Multi-parametric tissue characterization of brain neoplasms and their recurrence using pattern classification of MR images. *Acad Radiol* 15, pp. 966-977.
- [23] Østergaard L., Hochberg F.H., Rabinov J.D., Sorensen A.G., Lev M., Kim L., Weisskoff R.M., Gonzalez R.G., Gyldensted C., Rosen B.R., 1999. Early changes measured by magnetic resonance imaging in cerebral blood flow, blood volume, and blood-brain barrier permeability following dexamethasone treatment in patients with brain tumors. *J Neurosurg* 90, pp. 300-305.
- [24] Principi M., Italiani M., Guiducci A., Aprile I., Muti M., Giulianelli G., Ottoviano P., 2003. Perfusion MRI in the evaluation of the relationship between tumour growth, necrosis and angiogenesis in glioblastomas and grade 1 meningiomas. *Neuroradiology* 45, pp. 205-211.
- [25] Provenzale J.M., Mukundan S., Baroriak D.P., 2006. Diffusion-weighted and perfusion MR imaging for brain tumor characterization and assessment of treatment. *Radiology* 239, pp. 632-649.
- [26] Provenzale J.M., Mukundan S., Baroriak D.P., 2006. Diffusion-weighted and perfusion MR imaging for brain tumor characterization and assessment of treatment response. *Radiology* 239, pp. 632-649.
- [27] Provenzale J.M., Wang G.R., Brenner T., Petrella J.R., Sorensen A.G., 2002. Comparison of Permeability in High-Grade and Low-Grade Brain Tumors Using Dynamic Susceptibility Contrast MR Imaging. *Am J Roentgenol* 178, pp. 711-716.

- [28] Roberts H.C., Roberts T.P.L., Brasch R.C., Dillon W.P., 2000. Quantitative measurement of microvascular permeability in human brain tumors achieved using dynamic contrast – enhanced MR imaging: correlation with histologic grade. *Am J Neuroradiol* 21, pp. 891-899.
- [29] Roberts H.C., Roberts T.P.L., Lee T.Y., Dillon W.P., 2002. Dynamic Contrast – Enhanced CT of Human Brain Tumors: Quantitative Assessment of Blood Volume, Blood Flow, and Microvascular Permeability: Report of Two Cases. *Am J Neuroradiol* 23, pp. 828-832.
- [30] Siegal T., Rubinstein R.I., Tzuk-Shina T., Gomori J.M., 1997. Utility of relative cerebral blood volume mapping derived from perfusion magnetic resonance imaging in the routine follow up of brain tumors. *J Neurosurg* 86, pp. 22-27.
- [31] Smith S.M., Jenkinson M., Woolrich M.W., Beckmann C.F., Behrens T.E., Johansen-Berg H., Bannister P.R., De Luca M., Drobnjak I., Flitney D.E., Niazy R.K., Saunders J., Vickers J., Zhang Y., De Stefano N., Brady J.M., Matthews P.M., 2004. Advances in functional and structural MR image analysis and implementation as FSL. *Neuroimage* 23 (suppl 1).
- [32] Sugahara T., Korogi Y., Kochi M., Ikushima I., Hirai T., Okuda T., Shigematsu Y., Liang L., Ge Y., Ushio Y., Takahashi M., 1998. Correlation of MR Imaging – Determined Cerebral Blood Maps with Histologic and Angiographic Determination of Vascularity of Gliomas. *Am J Roengenol* 171, pp. 1479-1486.
- [33] Walecki J., Chojnacka E., 2007. Diagnostyka obrazowa guzów wewnątrzczaszkowych – część I – guzy neuroepitelialne. *Onkologia w praktyce klinicznej*, tom 3, nr 4, pp. 177-197.
- [34] Weber M.A., Zoubaa S., Schlieter M., Juttler E., Huttner H.B., Geletneky K., Ittrich C., Lichy M.P., Kroll A., Debus J., Giesel F.L., Hartmann M., Essig M., 2006. Diagnostic performance of spectroscopic and perfusion MRI for distinction of brain tumors. *Neurology* 66, pp. 1899-1906.
- [35] Young R.J., Knopp E.A., 2006. Brain MRI: tumor evaluation. *J Magn Reson Imaging* 24, pp. 709-724.
- [36] Zacharaki E.I., Wang S., Chawla S., Wolf R., Melhem E.R., Davatzikos C., 2009. Classification of brain tumor type and grade using MRI texture and shape in a machine learning scheme. *Magnetic Resonance in Medicine* 62, pp. 1609-1618.
- [37] Zimny A., Szaśiadek M., 2005. Badania perfuzyjne TK i MR – nowe narzędzie w diagnostyce guzów wewnątrzczaszkowych. *Adv Clin Exp Med*. 14; 3, pp. 583-592.

ANALIZA OBRAZU MRI U CHORYCH Z GUZEM OŚRODKOWEGO UKŁADU NERWOWEGO – PRÓBA OPRACOWANIA ALGORYTMU POSTĘPOWANIA

Streszczenie

Badanie metodą rezonansu magnetycznego jest aktualnie metodą referencyjną przy wykrywaniu i diagnozowaniu nowotworów centralnego układu nerwowego. Duża część nowotworów, zwłaszcza o wysokim stopniu złośliwości, charakteryzuje się większą zawartością wody w komórkach, co w badaniu MRI skutkuje wydłużeniem T1 i T2, uwidocznionym jako nasilenie sygnału w obrazach T2-zależnych oraz jego obniżeniem w obrazach T1-zależnych. MRI można przeprowadzić

z podaniem środka kontrastowego, co powoduje skrócenie czasu T1 i podniesienie sygnału w sekwencjach T1-zależnych. Pozwala to zidentyfikować obszary wzmożonej angiogenezy, która jest wykładnikiem stopnia złośliwości nowotworu oraz jego aktywności biologicznej. Otrzymane obrazy MRI są analizowane oraz oceniane przez radiologa, a następnie klinicystę. Najczęściej jest to analiza „na oko” i opiera się ona na ocenie obrazu MRI z ogólnie przyjętymi normami radiologicznymi. Jest to jednak metoda stosunkowo niedokładna, co sprawia, iż otrzymane obrazy MRI mogą zostać ocenione w sposób niewłaściwy, co z kolei może przyczynić się do postawienia złej diagnozy co do choroby pacjenta i wdrożenia lub wręcz brak wdrożenia odpowiedniego leczenia. Prowadzonych jest coraz więcej badań w zakresie wprowadzenia skomputeryzowanego algorytmu służącego do oceny badania MRI, jednak wypracowane metody są najczęściej bardzo skomplikowane i trudne do przeprowadzenia przez „laika” jakim jest klinicysta. Właśnie dlatego podjęta została próba opracowania w miarę prostego i czytelnego algorytmu analizy obrazu MRI u pacjentów z chorobą nowotworową centralnego układu nerwowego, która pozwoli na szybką i obiektywną ocenę badania rezonansu magnetycznego.

Słowa kluczowe: nowotwory, obrazowanie MRI, angiogeneza, analiza obrazu