SPECT Imaging of Pediatric Brain Tumor with Hexakis (Methoxyisobutylisonitrile) Technetium (I)

Lorcan A. O'Tuama, Alan B. Packard, and S. Ted Treves

Division of Nuclear Medicine, Department of Radiology, Children's Hospital, Harvard Medical School, Boston, Massachusetts

Technetium-99m-Hexamibi [Hexakis (methoxyisobutylisonitrile) technetium (I)] was developed as a myocardial perfusion agent with biologic properties similar to those of thallium-201 (201TI). As 201TI has recently been observed to be of value for the diagnosis of brain tumors when used in conjunction with single-photon emission computed tomography (SPECT) imaging technology, the possibility that the biologic similarity of the two radiopharmaceuticals extended to their affinity for tumors was tested. A 5-yr-old female patient with a brain stem astrocytoma showed marked focal uptake of ^{99m}Tc-Hexamibi at the site of tumor recurrence as defined by biopsy and prior ²⁰¹TI/SPECT study. Tumor-to-normal cortex radioactivity ratios for the ^{99m}Tc-Hexamibi/SPECT study were 132:1 and the spatial resolution of the 99mTc-Hexamibi images was high. This observation suggests that 99mTc-Hexamibi merits further study as a potential agent for SPECT imaging of human brain tumors.

J Nucl Med 1990; 31:2040-2041

hallium-201 (²⁰¹Tl), although developed for the evaluation of myocardial perfusion, has been found to be of value in the evaluation of tumor viability in adult and pediatric patients when used in conjunction with SPECT (1,2). The physical properties of technetium-99m (^{99m}Tc) (including gamma ray energy and radiation dose) are, however, superior to those of ²⁰¹Tl, and it would be preferable to have available a ^{99m}Tc radiopharmaceutical whose biologic properties paralleled those of ²⁰¹Tl. Several ^{99m}Tc radiopharmaceuticals have been developed as potential replacements for ²⁰¹Tl for the functional imaging of myocardial perfusion but none have been examined for possible uptake by tumors. One of the more promising classes of 99mTc myocardial perfusion agents is the hexakis (isonitrile) technetium (I) complexes (3,4), including hexakis (2methoxyisobutylisonitrile) technetium (I) (^{99m}Tc-Hexamibi) (Cardiolite, DuPont, N. Billerica, MA). This study reports the uptake of ^{99m}Tc-Hexamibi by a pediatric brain tumor.

CASE REPORT

The patient was a 5-yr-old girl who originally presented with right-sided clumsiness and hearing loss, and was treated by resection of a Grade 2 cerebellar astrocytoma followed by radiotherapy. Two years later she developed new neurologic deficits-increasing dysarthria, decrease in appetite, followed by a left peripheral facial palsy, and ataxia. Over the next 6-9 mo, a left gaze preference appeared. The computed tomography (CT) study showed increasing mass effect, but could not distinguish between recurrent tumor and radiation necrosis. At repeat posterior fossa craniectomy and C1 laminectomy, the right cerebellar hemisphere was very swollen, and the fourth ventricle was markedly displaced to the left. A frozen section of the cerebellum showed low-grade astrocytoma. Tumor was debulked in a plane extending through the cerebellar hemisphere up to the tentorium, and also at the brain stem level. Only subtotal removal could be achieved because of the diffuse intraaxial brainstem involvement. The patient was started on chemotherapy postoperatively. SPECT was performed 3 mo after the second resection. During this period, the patient showed increasing neurologic deficit. Magnetic resonance scans obtained 5 wk before, and 3 days after SPECT showed expansion of the brain stem and upper cord with increasing mass both within the cerebellar vermis and right hemisphere.

Informed consent was obtained prior to initiation of the study. The patient received 1.5 mCi of ²⁰¹Tl intravenously and data collection was started immediately postinjection. SPECT data were collected in 64 steps of 30 sec each through 360° using a rotating gamma camera (Siemens Orbiter 7500, Hoffman Estates, IL) equipped with a 30° slant-hole collimator. The patient then received 10 mCi of ^{99m}Tc-Hexamibi and a second data set was collected using the same conditions as for ²⁰¹Tl SPECT. The data were processed and image reconstruction performed with an ADAC 3300 system using a Butterworth filter and filtered backprojection (ADAC Laboratories, Milpitas, CA). Tumor-to-normal tissue ratios were calculated using the average count rate per pixel in a region of interest including the tumor divided by the average count rate per pixel for a region in the contralateral cerebral cortex.

Received Apr. 2, 1990; revision accepted Jun. 7, 1990.

For reprints contact: L.A. O'Tuama, MD, Division of Nuclear Medicine, Children's Hospital, 300 Longwood Ave., Boston, MA 02115.

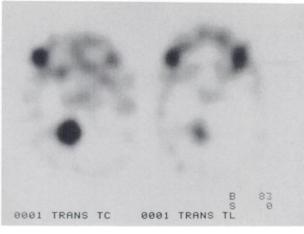


FIGURE 1

Technetium-99m-Hexamibi (left panel) and ²⁰¹TI (right panel) SPECT images in the transaxial plane with image contrast adjusted for optimal contrast in the ²⁰¹TI image.

Focal uptake of ^{99m}Tc-Hexamibi was noted involving the tumor site in the right anterior cerebellum (Fig. 1). There was minimal uptake of the tracer by normal brain. Technetium-99m-Hexamibi showed greater uptake at the tumor site than ²⁰¹Tl, and defined the margins of the lesion more clearly. Tumor-to-normal cortex ratios were 132:1 for ^{99m}Tc-Hexamibi and 80:1 for ²⁰¹Tl.

DISCUSSION

The area of focal uptake of ^{99m}Tc-Hexamibi corresponds to active growth of recurrent tumor as demonstrated by the ²⁰¹Tl study as well as recent biopsy and clinical evidence of active disease. Visual inspection indicates and quantitative analysis confirms that ^{99m}Tc-Hexamibi is taken up to a greater extent by the tumor than is ²⁰¹Tl. This difference is illustrated by the different gray scales required to optimize the ^{99m}Tc-Hexamibi and ²⁰¹Tl images (Fig. 1). The distribution of both tracers within the tumor is similar.

It is particularly interesting to note that the tumorto-normal cortex ratios observed in the ^{99m}Tc-Hexamibi/SPECT study not only exceed those observed with 201 Tl by more than 50% but also exceed by almost twentyfold the values reported for PET studies of brain tumors using [11 C]L-methionine, the most tumor-avid, positron-emitting radiopharmaceutical that has been developed to date (5).

Studies are currently underway to determine if similar results are observed with tumors other than pediatric astrocytomas as well as to elucidate the mechanism of uptake of ^{99m}Tc-Hexamibi by the tumor. If a similar pattern of uptake is observed for other types of tumors, ^{99m}Tc-Hexamibi has the potential to offer significant clinical advantages over both ²⁰¹Tl SPECT and PET while incorporating the practical advantages inherent in a kit-based ^{99m}Tc radiopharmaceutical.

ACKNOWLEDGMENTS

We thank Stephen Sallan, MD, R. Michael Scott, MD, and Nancy Tarbell, MD, for referring the patient for the SPECT study. James S. Ulanski, CNMT provided meticulous technical assistance.

REFERENCES

- Black KL, Randall AH, Kim KT, Becker DP, Lerner C, Marciano D. Use of thallium-201 SPECT to quantitate malignancy grade of gliomas. J Neurosurg 1989; 71:342-346.
- Ulanski JS, O'Tuama LA, Davis RT, et al. Pediatric brain tumor imaging with ²⁰¹Tl and ^{99m}Tc-HMPAO [Abstract]. J Nucl Med Technol 1990; 18:140.
- 3. Piwnica-Worms D, Kronauge JF, Holman BL, et al. Hexakis (carbomethoxyisopropylisonitrile) technetium (I), a new myocardial perfusion imaging agent: binding characteristics in cultured chick heart cells. J Nucl Med 1988; 29:55-61.
- Wackers FJTh, Berman DS, Maddahi J, et al. Technetium-99m-hexakis 2-methoxyisobutyl isonitrile: human biodistribution, dosimetry, safety, and preliminary comparison to thallium-201 for myocardial perfusion imaging. J Nucl Med 1989; 30:301-311.
- Bergstroem M, Collins VP, Ehrin E, et al. Discrepancies in brain tumor extent as shown by computed tomography and positron emission tomography using (gallium-68) EDTA, ¹¹Cglucose and ¹¹C-methionine. J Comput Assist Tomogr 1983; 7:1062–1066.