

BRAIN SPECT IN THE PRE-SURGICAL EVALUATION OF EPILEPTIC PATIENTS

PRELIMINARY RESULTS

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SUMMARY — Pre-surgical evaluation of epileptic patients consists of neurological examination, intensive electroencephalographic (EEG) monitoring and anatomical studies (CT and MRI). Functional methods such as PET and SPECT imaging are now used more frequently. We have studied pre-operatively 15 adult epileptic patients (8 female, 7 male) using a rotational scintillation camera interfaced to a dedicated computer. The tomographic images were obtained 15 minutes after intravenous injection of ^{99m}Tc -HMPAO. All had MRI scanning and intensive EEG monitoring which generally included seizure recording. Five patients had progressive lesions (3 meningiomas, 2 astrocytomas). In 10 patients, neuroradiological studies did not show the presence of progressive lesions (2 normal scans and 8 cases with inactive lesions). Two patients with meningioma showed hypoperfusion at the lesion site while the third patient had a marked hyperperfusion which might correlate with the clinical diagnosis of *epilepsia partialis continua*. In the astrocytoma patients SPECT scans showed hypoperfusion at the lesion site. Data obtained from the 10 patients without progressive CNS lesions showed: (a) in 4, SPECT findings correlated well with the anatomical findings; (b) in 5 instances, SPECT was able to disclose additional functional deficits; (c) in one case, there was no SPECT correlate of a discrete anatomical lesion. In 5 of these cases with no progressive lesions (n=10) SPECT findings were useful as a complementary tool in determining the clinical or surgical management of these patients. Despite the small number and heterogeneity of the present sample, SPECT seems to be an useful tool as part of the clinical workup of epileptic patients who are candidates for epilepsy surgery.

KEY WORDS: SPECT, epilepsy, surgery, pre-operative evaluation.

SPECT cerebral na avaliação pré-cirúrgica de pacientes epilépticos

RESUMO — A avaliação pré-operatória de pacientes epilépticos consiste basicamente de exame neurológico, monitorização intensiva de EEG e estudos anatômicos (CT e RMN). Métodos funcionais como PET e SPECT têm sido utilizados mais frequentemente nos últimos anos. Foi realizada a investigação pré-operatória em 15 pacientes epilépticos adultos (8 mulheres, 7 homens) utilizando câmara de cintilação rotacional acoplada a computador. As imagens tomográficas foram obtidas 15 minutos após a injeção de ^{99m}Tc HMPAO. Todos os pacientes foram submetidos a RMN e monitorização intensiva de EEG, que geralmente inclui registros ictais. Cinco pacientes possuíam lesões expansivas (3 meningiomas e 2 astrocitomas). Em 10 pacientes, os estudos neurorradiológicos não mostraram presença de lesões expansivas (2 normais e 8 com lesões inativas). O estudo de 2 pacientes com meningiomas mostrou presença de hipoperfusão ao nível da lesão, enquanto em um terceiro caso marcada hiperperfusão foi detectada e, neste paciente, provavelmente correlacionava-se quadro de *epilepsia partialis continua*. Nos pacientes com astrocitomas, os exames de SPECT mostraram hipoperfusão no local da lesão. Os dados obtidos em pacientes não portadores de lesões expansivas do sistema nervoso mostraram: (a) em 4 casos os achados de SPECT correlacionaram-se bem aos achados

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anatômicos; (b) em 5 casos, os achados de SPECT mostraram déficits de fluxo adicionais a lesão anatômica; (c) em um caso, não houve correlação entre os achados de SPECT e uma pequena lesão anatômica. Em 5 casos com lesões não progressivas (n=10) os achados de SPECT foram úteis na determinação da conduta clínica ou cirúrgica. Apesar do pequeno número e da heterogeneidade da presente amostra, o SPECT parece ser importante método complementar no arsenal clínico-propedêutico de pacientes epiléticos candidatos a cirurgia.

PALAVRAS CHAVES: SPECT, epilepsia, cirurgia, avaliação pré-operatória.

Epilepsy is a chronic neurologic disease characterized by recurrent ictal episodes derived from the synchronous hyperactivity of cortical neuronal populations. The main clinical workup of epileptic patients comprises a seizure history, neurological examination, electroencephalographic (EEG) recordings and neuroradiological studies (CT and MRI). 15-20% of the patients with epilepsy are medically refractory to the current anti-epileptic drugs regimens and are potential candidates for epilepsy surgery. In this selected population, accurate lateralization and localization of the epileptogenic region is mandatory. Functional methods (PET and SPECT) have been used in an increasing frequency as a tool in the pre-surgical evaluation of epileptic patients in the last years^{2,3}. PET and SPECT studies have shown that the epileptic regions were characterized by hypoperfusion/hypometabolism (hypo/hypo) patterns in the interictal period, even when there was no anatomical correlate in the neuroradiological studies. This hypo/hypo region is often bigger than the actual epileptic focus⁶⁻⁸.

Our preliminary SPECT (single photon emission computerized tomography) scanning results obtained from adult epileptic patients who were candidates for epilepsy surgery are described in this paper.

METHODS

Fifteen adult patients (eight males, seven females) who were candidates for epilepsy surgery were submitted to interictal SPECT scanning. One patient had an ictal and another one a post-ictal study. Five patients had progressive central nervous system (CNS) diseases (three meningiomas and two astrocytomas). In ten patients the neuroradiological studies did not disclose any progressive lesion (two normal and eight with inactive lesions scans) (Table 1).

The brain SPECT images were obtained after intravenous injection of 740 MBq (20 mCi) of ^{99m}Tc-hexamethylenpropyloximeamine (HMPAO) (Ceretek/Amersham). The vials were reconstituted using 5 ml of ^{99m}Tc-pertechnetate solution with activity not superior to 1295 MBq (35 mCi). Radiochemical controls (mean 86%) were obtained. All patients received 500 mg of potassium perchlorate 30 minutes prior to the injection. Visual, sensitive or auditive stimuli were avoided during radiopharmaceutical administration.

Tomographic images were obtained 15 minutes after injection, using a rotational single-head scintillation camera (Siemens model Orbiter) interfaced to a dedicated computer (Microdelta) and equipped with a high resolution colimator. The image data were acquired for 30 seconds per stop, for a total of 64 stops, using a 360 degrees rotation over the circumference of the head. The matrix size used was 64 x 64 and a proper center of rotation and flood uniformity corrections were applied. Transaxial, sagittal and coronal slices were obtained by processing procedures by means of a Butterworth filter. The slice size was 1 pixel or 6 mm thick.

RESULTS

SPECT data can be summarized as in the Table 1. Two patients with meningioma showed hypoperfusion at the lesion site while the third patient had a marked hyperperfusion which might correlate with the clinical diagnosis of epilepsia partialis continua (Fig. 1).

In the patients with astrocytoma, SPECT scans showed hypoperfusion at the lesion site. Data obtained from the ten patients without progressive CNS lesions showed: (a) in four, SPECT findings correlated well with the anatomical findings; (b) in five instances, SPECT was able to disclose additional functional deficits (Fig. 2); (c) in one case, there was no SPECT correlate of a discrete anatomical lesion. In five of these cases with no progressive lesions (n=10) SPECT findings were useful as a complimentary tool in determining the clinical or surgical management of these patients. In one case a post-ictal SPECT study disclosed an enlarged hypoperfused area when compared to a previous inter-ictal exam (Fig. 3).

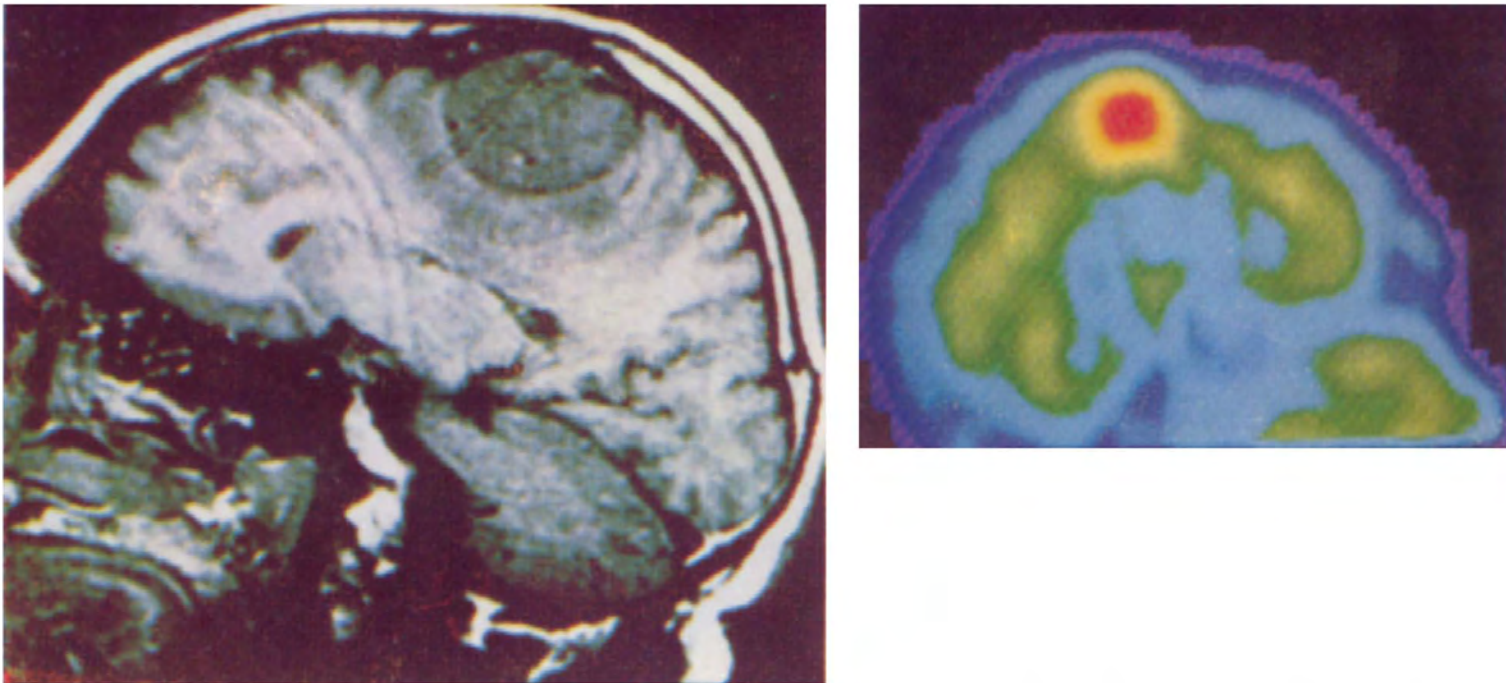


Fig. 1 — Sagittal MRI (left) and ictal SPECT (right) images in a patient with meningioma showing a hyperperfusion pattern.

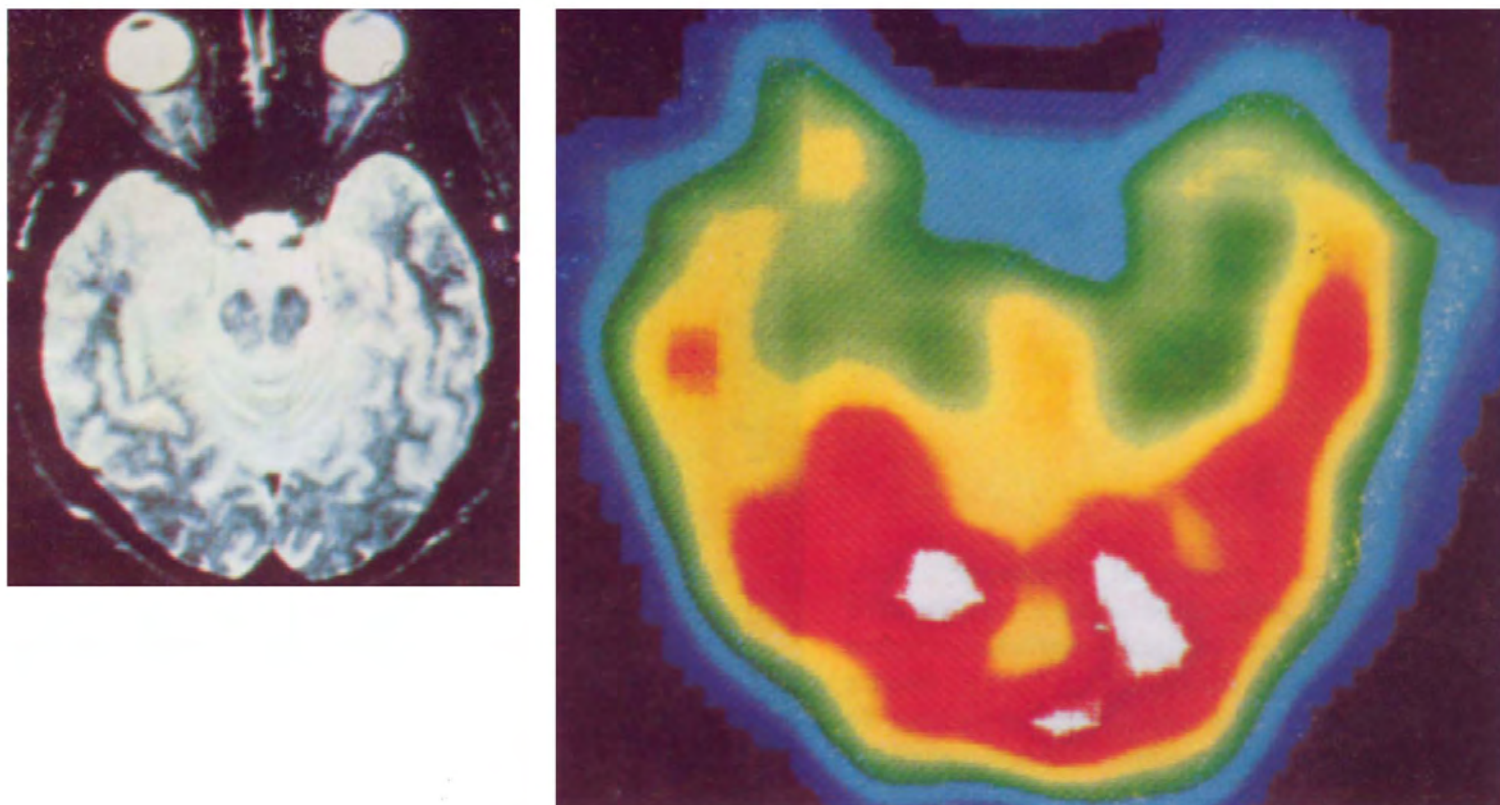


Fig. 2 — Right temporal lobe hypoperfusion in a patient with a normal MRI. Normal transverse MRI slice (left) and abnormal interictal SPECT study (right).

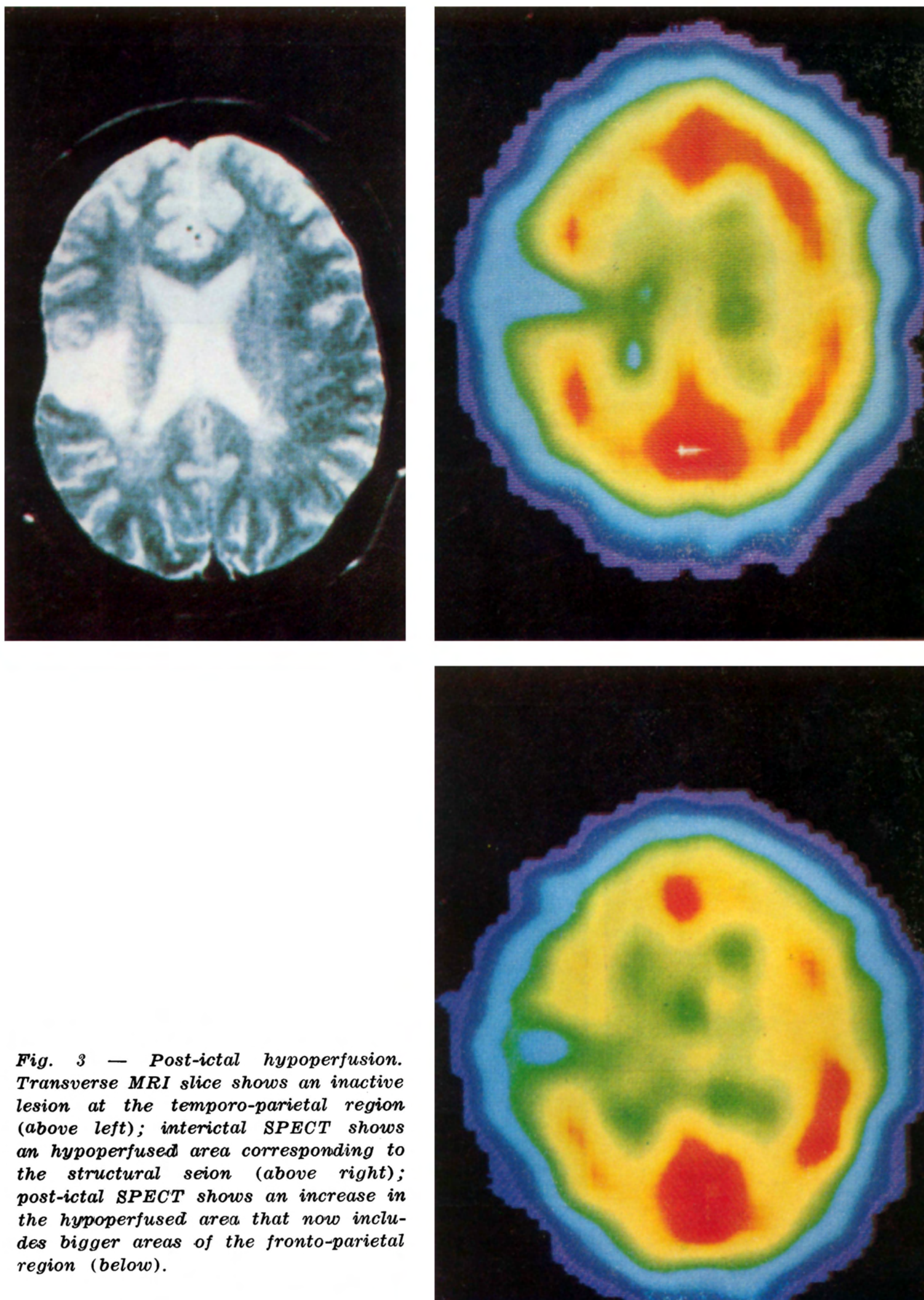


Table 1 — Clinical and imaging data.

| Pt | Clinical | EEG | MRI | SPECT |
|----|----------|------------|------------------------|---|
| 1 | SPS | RF | RF Meningioma | RF Hypoperfusion |
| 2 | SPS | RF, RO, LO | RO Inactive lesion | RF, TPO, LO Hypoperfusion |
| 3 | SPS | RT | Normal | RT Hypoperfusion |
| 4 | CPC | RT, LT | LT Astrocytoma | LT Hypoperfusion |
| 5 | | RF | RF Inactive lesion | RF, LC Hypoperfusion |
| 6 | CPC | LT | LT Inactive lesion | -Inter-ictal: LTP Hypoperfusion -Post-ictal LTP more intense hypoperfusion |
| 7 | SPS | LFP | LFP Meningioma | LFP Hypoperfusion |
| 8 | CPC | LF, RT | LF Inactive lesion | LF Hypoperfusion |
| 9 | SPS | LFTPO | L Cerebral hemiatrophy | LT Hypoperfusion |
| 10 | CPC | LT, RT | LT Inactive lesion | LF Hypoperfusion |
| 11 | CPC | LT | Normal | LT Hypoperfusion |
| 12 | CPC | RT | RT Meningioma | RTPO Hypoperfusion |
| 13 | CPC | RT, LT | RT Astrocytoma | RT Hypoperfusion |
| 14 | CPC | RT | RT Dysgenesis | Normal |
| 15 | SPS | LF | LF Dysgenesis | LF Hypoperfusion |

Pt, patients; Clinical, seizure pattern; SPS, single partial seizure; CPS, complex partial seizure; F, frontal; P, parietal; T, temporal; O, occipital; C, cerebellum; L, left; R, right.

COMMENTS

Epilepsy is a disease caused by the overactivity of neuronal populations. On the other hand, functional methods have shown that the epileptic focus is characterized by a hypo/hypo pattern in the interictal state. This would suggest an apparent lack of consistency between the actual pathophysiology of epilepsy and the results obtained from functional studies. The fact that the epileptogenic region is a lesionated area should be taken into account and should explain the previous findings⁹. The hypo/hypo region seen in these cases is often larger than the actual epileptogenic area¹. Cortico-cortical inhibition in the periphery of the focus and an expanded area of functional derangement surrounding the epileptogenic foci might be involved in this process. On the contrary, ictal patterns were characterized by an hyperperfusion/hypermetabolic pattern (hyper/hyper) and correlated well to a marked synchronization and hyperactivity of large neuronal populations⁴. Post-ictal studies have shown a marked and/or enlarged hypo/hypo pattern in the epileptic focus. Studies performed immediately after the ictus might show hyper/hyper zones surrounded by hypo/hypo ones⁵.

Our studied population was heterogeneous. Five cases had progressive lesions. Such cases were commonly associated to brain edema. The analysis of functional data obtained from SPECT in such patients was difficult and should be considered carefully. On the other hand, the analysis of the cases with no progressive lesions (n=10) yielded a high diagnostic rate.

Ictal studies are more difficult to obtain than interictal ones. In this series, only one patient with *epilepsia partialis continua* associated to a meningioma had an ictal SPECT which showed a marked hyper/hyper pattern.

Post-ictal studies are more easily obtained but their results can vary widely according to the timing of the injection of the tracer. In case 6, an enlargement of the interictal hypo/hypo previously determined area was seen post-ictally.

In five cases with no progressive lesions (n=10) SPECT was an important item in the presurgical management protocol and was used to determine a medical and/or surgical procedure.

Despite the fact that PET scanning yields a better image resolution than SPECT, the latter has been being used in an increasing frequency in the last years. It is an easier method and certainly a less expensive one. The above data suggest that SPECT is an useful tool in the presurgical evaluation of epileptic patients. Homogeneous population of patients with unilateral or bilateral temporal foci with normal MRI are now being studied in our Center.

REFERENCES

1. Abou-Khalil BW, Siegel GJ, Sackellares JC. PET studies of cerebral glucose metabolism in chronic partial epilepsies. *Ann Neurol* 1987, 22:480-486.
2. Engel JJr, Babb TL, Phelps ME. Contribution of positron emission tomography to understanding mechanisms of epilepsy. In Engel J (ed): *Fundamental Mechanisms of Human Brain Function*. New York: Raven Press, 1987, p 209-218.
3. Lee BI, Markand ON, Sidigui AR, Park HM, Mock B, Wellman HH, Worth RM, Edwards MK. SPECT brain imaging using HIPDM: intractable complex partial seizures. *Neurology* 1986, 36:1471-1477.
4. Ochs RF, Gloor P, Tyler JF. Effects of generalized spike-and-wave discharge on glucose metabolism measured by PET. *Ann Neurol* 1987, 21:458-464.
5. Sackellares JC, Siegel JC, Abou-Khalil BW, Hood TW, Gilman S, McKeever PE, Hichwa RD, Hutchins GD. Differences between lateral and mesial temporal metabolism interictally in epilepsy of mesial temporal origin. *Neurology* 1990, 40:1420-1426.
6. Stefan H, Bocher-Schwarz HG, Biersack HJ, Burr W, Penin H, Heiss WD. Functional and morphological abnormalities in temporal lobe epilepsy: a comparison of interictal and ictal EEG, CT, MRI, SPECT and PET. *J Neurol* 1987, 234:377-384.
7. Swartz BE, Halgren E, Delgado-Escueta AV, Mandelkern M, Gee M, Quinones N, Bland WH, Repchan J. Neuroimaging in patients with seizures of probable frontal lobe origin. *Epilepsia* 1989, 30:547-558.
8. Theodore WH, Dorwart R, Holmes M. Neuroimaging in refractory partial seizures: comparison of PET, CT and MRI. *Neurology* 1986, 36:750-759.
9. Theodore WH, Fishbein D, Dubinsk R. Patterns of cerebral glucose metabolism in patients with partial seizures. *Neurology* 1988, 38:1201-1206.