

Usefulness of technetium-99m hexamethylpropylene amine oxime lung scan to detect inhalation lung injury of patients with pulmonary symptoms/signs but negative chest radiograph and pulmonary function test findings after a fire accident—a preliminary report

Yu-Chien SHIAU,* Feng-Yuan LIU,* Jeffrey J.P. TSAI,** Jhi-Joung WANG,***
Shung-Tai HO**** and Albert KAO*****

*Department of Nuclear Medicine, Far Eastern Memorial Hospital, Taipei;

**Graduate Institute of Bioinformatics, Taichung Healthcare and Management University, Taichung;

***Department of Medical Research, Chi-Mei Medical Center, Tainan;

****School of Medicine, National Defense Medical Center, Taipei;

*****Department of Medical Research, China Medical University Hospital, Taichung; Taiwan

Objective: In this study, we employed technetium-99m hexamethylpropylene amine oxime (^{99m}Tc HMPAO) lung scan to detect inhalation lung injury of patients after a fire accident. **Methods:** Ten healthy men for controls and 10 male patients with pulmonary symptoms/signs from a fire accident were enrolled in this study for comparison. ^{99m}Tc HMPAO lung scan was performed in each control and patient, as well as the degree of pulmonary vascular endothelium damage was represented as lung/liver uptake ratios (L/L ratio). All of the controls and patients had no smoking histories. None of the controls and patients had positive findings of plain chest radiograph (CXR) and pulmonary function test (PFT). **Results:** The results showed that significantly higher L/L ratio in the 10 patients (0.53 ± 0.07) than in the 10 controls (0.30 ± 0.07) (the p value < 0.05). Using a cut-off value of 0.40, all of the 10 patients had abnormally increased L/L ratios. **Conclusions:** We conclude that ^{99m}Tc HMPAO lung scan has the potential to be a sensitive, objective and noninvasive method to detect inhalation lung injury of patients with pulmonary symptoms/signs but negative CXR and PFT findings after a fire accident.

Key words: technetium-99m hexamethylpropylene amine oxime, inhalation lung injury, fire accident

INTRODUCTION

SMOKE OR HEAT INHALATION after a fire accident usually causes parenchymal lung injury results in interference with gas exchange.^{1–3} This often leads to acute pulmonary insufficiency, which is a major cause of morbidity and mortality in fire victims.^{1–3} Early reports suggested that the plain chest radiograph (CXR) and pulmonary function

test (PFT) were of little value in the early diagnosis and management of patients with inhalation lung injury after a fire accident.^{4–6} However, there is still a lack of the sensitive, objective and noninvasive method to detect inhalation lung injury of patients from a fire accident. Technetium-99m hexamethylpropylene amine oxime (^{99m}Tc HMPAO) lung scan has been used to assess mild lung injury in various diffuse infiltrative lung disease, which are presented as the degree of pulmonary vascular endothelium damage.^{7,8} Therefore, in this study, we tried to use ^{99m}Tc HMPAO lung scan to detect inhalation lung injury of patients with pulmonary symptoms/signs, since either chest radiograph (CXR) or pulmonary function test (PFT) resulted in little value after a fire accident.

Received January 20, 2003, revision accepted March 10, 2003.

For reprint contact: Albert Kao, M.D., Department of Medical Research, China Medical University Hospital, No. 2, Yuh-Der Road, Taichung 404, TAIWAN.

E-mail: albertkaotw@yahoo.com.tw.

MATERIALS AND METHODS

Subjects

Ten male patients (aged from 30–49 years old, mean: 40.6 ± 6.3 years old) with pulmonary symptoms/signs such as dyspnea and/or cough with/without sputum following a fire accident in a supermarket were sent to the emergency room for further evaluation. However, to estimate the amount of inhaled smoke and materials of combusted in the fire is difficult. Patients with a history of smoking or previous lung disease were excluded. No patient's occupation was firefighter or daily exposure to smoke. In addition, 10 healthy men without smoking histories were included as normal controls. All of the study subjects including controls and patients had negative CXR and

PFT findings. Negative CXR findings excluded pleurisy, atelectasis, pulmonary edema, acute pneumonitis, diffuse interstitial disease and diaphragmatic dysfunction with loss of lung volume.⁹ Comprehensive PFT included forced vital capacity (FVC), forced expiratory volume in one second (FEV₁), and diffusion capacity by a single breath CO method (DL_{co}). Above the 80 percent of the predicted value was considered negative findings. ^{99m}Tc HMPAO lung scan, CXR, and PFT were performed within 24 hours. The informed consent was obtained for the participating patients.

^{99m}Tc HMPAO Lung Scan

Anterior planar image of ^{99m}Tc HMPAO lung scan including all of the lungs and most of the liver was obtained

Table 1 Detailed data of controls and patients

Case No.	Controls		Patients			
	Age (years)	Lung/Liver ratio	Age (years)	Lung/Liver ratio		Hospitalization /Days
1	30	0.24	31	0.46	*	No
2	34	0.32	33	0.66	*	Yes/2
3	35	0.38	34	0.48	*	No
4	37	0.39	35	0.49	*	No
5	40	0.19	39	0.54	*	Yes/3
6	44	0.34	41	0.52	*	No
7	45	0.26	42	0.42		No
8	46	0.23	43	0.62	*	Yes/3
9	46	0.31	45	0.50	*	No
10	49	0.37	48	0.55	*	Yes/2
Max.	49	0.39	48	0.66		
Min.	30	0.19	31	0.42		
Mean	40.6	0.3	39.1	0.53		
SD	6.3	0.07	5.6	0.07		

*: abnormal results, ratio > 0.30 + 0.07 × 2 = 0.44

Table 2 Detailed data of pulmonary function tests in controls and patients

Case No.	Controls			Symptoms & Signs	Patients		
	FVC % predicted	FEV ₁ % FVC	DL _{co} %		FVC % predicted	FEV ₁ % FVC	DL _{co} %
1	91	88	105	dyspnea, cough	94	83	104
2	90	91	110	cough, dyspnea	108	90	112
3	110	85	95	cough, sputum	91	83	93
4	103	93	90	cough, sputum	98	94	95
5	88	87	98	cough, sputum	102	99	105
6	96	96	103	dyspnea, sputum	93	91	99
7	97	93	121	cough	90	85	101
8	95	89	93	cough, sputum	110	97	109
9	92	90	107	cough, dyspnea	91	89	96
10	105	94	102	dyspnea, cough	101	92	90
Max.	110	96	121		110	99	112
Min.	88	85	90		90	83	90
Mean	96.7	90.6	102		97.8	90.3	100
SD	7.2	3.4	9.1		7.2	5.5	7.1

10 minutes after intravenous injection of 740–925 MBq (20–25 mCi) of ^{99m}Tc HMPAO. ^{99m}Tc HMPAO lung scans were acquired using a gamma camera interfaced with a computer. Analog images and computerized images with an accumulation of 500 k counts were obtained. Regions-of-interests (ROI) were drawn over the mid-portion of the right lung and the highest activity area of the liver parenchyma was selected and lung/liver uptake ratios (L/L ratio) were calculated to represent the degree of pulmonary vascular endothelium damage based on the published method.⁸

Statistical Analysis

The statistical difference of mean lung/liver uptake ratio between the 10 controls and 10 patients was analyzed by a nonparametric Mann-Whitney U test. If the p value < 0.05, the statistical difference was considered as significant.

RESULTS

Detailed data of lung/liver uptake ratio in controls and patients are shown in Table 1. Detailed data of pulmonary function tests in controls and patients included FVC, FEV₁, and DLco were listed in Table 2. The L/L ratio was 0.30 ± 0.07 in the 10 normal controls and 0.53 ± 0.07 in the 10 patients. The difference was significant with a p value of less than 0.05. Using a cut-off value of 0.40 (all of 10 normal controls had L/L ratios below 0.39), all of the 10 (100%) patients had an abnormally increased L/L ratio. Four patients were admitted for further treatments with 2–3 days of hospitalization. The L/L ratio of the 4 patients with hospitalization was higher (0.59 ± 0.06) than those of the 6 patients without hospitalization (0.48 ± 0.03). However, the difference is not significant (p value > 0.05). Following-up of 3 months after the 10 patients were discharged, no patient had pulmonary symptoms/signs and sequels.

DISCUSSION

^{99m}Tc HMPAO is a cyclic amine as a lipophilic brain imaging agent used for the diagnosis of stroke and dementias. Elevated pulmonary uptake of this agent has been demonstrated in patients with pulmonary edema¹⁰ and in smokers.¹¹ Its localization is presumably in the pulmonary vascular endothelium.¹² However, in this study, the mechanism of increased uptake of ^{99m}Tc HMPAO in the lungs of patients with pulmonary symptoms/signs but negative CXR and PFT findings after a fire accident is unknown. It may be related to dysfunction of vascular endothelial cells or dysfunction of amine metabolism in the injured endothelium.¹³

The value of the CXR in the evaluation of the inhalation lung injury in fire victims is debatable. Previous studies suggested that the value of CXR in the diagnosis and management of patients after acute smoke inhalation is

limited.^{4–6} In our study, only the patients with negative CXR findings were included. Therefore, we conclude that CXR is not a sensitive modality to detect mild inhalation lung injury after a fire accident. Abnormality in CXR may not be seen until a severe complication such as atelectasis, alveolar edema, or bronchopneumonia develops. Because only PFT after fire accident could be done in this study, assessment of the influence of inhalation lung injury on PFT due to a fire accident is very difficult. However, compare with a normal database, only the patients with negative PFT findings (above 80% of normal predicted values) were included in this study. Therefore, our preliminary results suggest that PFT may not be a sensitive test to detect mild inhalation lung injury after a fire accident.

Although the number of patients in our study was too small to make a conclusion, our findings show significantly increased L/L ratios in patients with pulmonary symptoms/signs but negative CXR and PFT findings after a fire accident. We have to consider that the conventional CXR and PFT are not sensitive modalities to detect mild inhalation lung injury after a fire accident. In conclusion, ratio of lung/liver uptake of ^{99m}Tc HMPAO served as an additional or alternative way, independent of CXR and PFT seems to be a sensitive, objective and noninvasive method to early detect mild inhalation lung injury of patients after a fire accident. However, future investigations such as broncho-alveolar lavage and cytological examination in a larger series of case numbers with longer follow-up and their outcomes should be encouraged to confirm our findings.

REFERENCES

1. Fein A, Leff A, Hopewell PC. Pathophysiology and management of the complications resulting from fire and the inhaled products of combustion. *Crit Care Med* 1980; 8: 94–98.
2. Peitzman AB, Shires GT III, Corbett WA, Curreri PW, Shires GT. Measurement of lung water in inhalation injury. *Surgery* 1981; 90: 305–312.
3. Kimmel EC, Still KR. Acute lung injury, acute respiratory distress syndrome and inhalation injury: an overview. *Drug Chem Toxicol* 1999; 22: 91–128.
4. Eaton RJ, Senior RM, Pierce JA. Aspects of respiratory care pertinent to the radiologist. *Radio Clin N Amer* 1973; 2: 93–107.
5. Lin WY, Kao CH, Wang SJ. Detection of acute inhalation injury in fire victims by means of technetium-99m DTPA radioaerosol inhalation lung scintigraphy. *Eur J Nucl Med* 1997; 24: 125–129.
6. Wittram C, Kenny JB. The admission chest radiograph after acute inhalation injury and burn. *Br J Radiol* 1994; 67: 751–754.
7. Niden AH, Mishkin FS. Diffuse infiltrative lung disease. In: *Pulmonary nuclear medicine—techniques in diagnosis of lung disease*, Atkins HL (ed), New York; Marcel Dekker Inc., 1984: 167–200.

8. Shih CM, Shiau YC, Wang JJ, Ho ST, Kao A. Increased lung uptake of technetium-99m hexamethylpropylene amine oxime in systemic lupus erythematosus. *Respiration* 2002; 69: 143–147.
9. Tocino IM. Chest radiology and other imaging techniques. In: *The lung in rheumatic disease*, Cannon GW, Zimmerman GA (eds), New York; Marcel Dekker Inc., 1990: 81–116.
10. Owada K, Uwantono E, Takeda H. Investigation of myocardial and pulmonary uptake of Tc-99m-HMPAO in the patients with cardiac diseases (abstract). *KAKU IGAKU (Jpn J Nucl Med)* 1993; 30: 960.
11. Shih WJ, Gruenwald F, Biersack HJ, Berger R, Brandenburg S, Coupal J, et al. Tc-99m HMPAO diffuse pulmonary uptake demonstrated in cigarette smokers. *Clin Nucl Med* 1991; 16: 668–672.
12. Touya JJ, Rahimian J, Grubbs DE, Corbus HF, Bennett LR. A noninvasive procedure for *in vivo* assay of a lung amine endothelial receptor. *J Nucl Med* 1985; 26: 1302–1307.
13. Suga K, Uchisako H, Nishigauchi K, Shimizu K, Kume N, Yamada N, et al. Technetium-99m-HMPAO as a marker of chemical and irradiation lung injury: experimental and clinical investigations. *J Nucl Med* 1994; 35: 1520–1527.