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#### Contents

#### Thrice Monthly Volume 10 Number 2 January 14, 2022

#### **EDITORIAL**

397 New trends in treatment of muscle fatigue throughout rehabilitation of elderlies with motor neuron diseases

Mohamed A

#### **MINIREVIEWS**

- 401 What emotion dimensions can affect working memory performance in healthy adults? A review Hou TY, Cai WP
- 412 Quadrilateral plate fractures of the acetabulum: Classification, approach, implant therapy and related research progress

Zhou XF, Gu SC, Zhu WB, Yang JZ, Xu L, Fang SY

#### **ORIGINAL ARTICLE**

#### **Case Control Study**

Methylprednisolone accelerate chest computed tomography absorption in COVID-19: A three-centered 426 retrospective case control study from China

Lin L, Xue D, Chen JH, Wei QY, Huang ZH

#### **Retrospective Study**

437 Analysis of photostimulable phosphor image plate artifacts and their prevalence Elkhateeb SM, Aloyouny AY, Omer MMS, Mansour SM

448 N6-methyladenine-modified DNA was decreased in Alzheimer's disease patients Lv S, Zhou X, Li YM, Yang T, Zhang SJ, Wang Y, Jia SH, Peng DT

458 Inflammation-related indicators to distinguish between gastric stromal tumors and leiomyomas: A retrospective study

Zhai YH, Zheng Z, Deng W, Yin J, Bai ZG, Liu XY, Zhang J, Zhang ZT

469 Relationship between Ki-67 and CD44 expression and microvascular formation in gastric stromal tumor tissues

Ma B, Huang XT, Zou GJ, Hou WY, Du XH

477 Modified surgical method of supra- and infratentorial epidural hematoma and the related anatomical study of the squamous part of the occipital bone

Li RC, Guo SW, Liang C

485 Combined molybdenum target X-ray and magnetic resonance imaging examinations improve breast cancer diagnostic efficacy

Gu WQ, Cai SM, Liu WD, Zhang Q, Shi Y, Du LJ



World Journal of Clinical Cases		
Conter	nts Thrice Monthly Volume 10 Number 2 January 14, 2022	
492	Value of thyroglobulin combined with ultrasound-guided fine-needle aspiration cytology for diagnosis of lymph node metastasis of thyroid carcinoma	
	Zhang LY, Chen Y, Ao YZ	
502	Locking compression plate + T-type steel plate for postoperative weight bearing and functional recovery in complex tibial plateau fractures	
	Li HF, Yu T, Zhu XF, Wang H, Zhang YQ	
511	Effect of Mirena placement on reproductive hormone levels at different time intervals after artificial abortion	
	Jin XX, Sun L, Lai XL, Li J, Liang ML, Ma X	
518	Diagnostic value of artificial intelligence automatic detection systems for breast BI-RADS 4 nodules	
	Lyu SY, Zhang Y, Zhang MW, Zhang BS, Gao LB, Bai LT, Wang J	
	Clinical Trials Study	
528	Analysis of 20 patients with laparoscopic extended right colectomy	
	Zheng HD, Xu JH, Liu YR, Sun YF	
	Observational Study	
538	Knowledge, attitude, practice and factors that influence the awareness of college students with regards to breast cancer	
	Zhang QN, Lu HX	
547	Diagnosing early scar pregnancy in the lower uterine segment after cesarean section by intracavitary ultrasound	
	Cheng XL, Cao XY, Wang XQ, Lin HL, Fang JC, Wang L	
554	Impact of failure mode and effects analysis-based emergency management on the effectiveness of craniocerebral injury treatment	
	Shao XL, Wang YZ, Chen XH, Ding WJ	
563	Predictive value of alarm symptoms in Rome IV irritable bowel syndrome: A multicenter cross-sectional study	
	Yang Q, Wei ZC, Liu N, Pan YL, Jiang XS, Tantai XX, Yang Q, Yang J, Wang JJ, Shang L, Lin Q, Xiao CL, Wang JH	
	Prospective Study	
576	5-min mindfulness audio induction alleviates psychological distress and sleep disorders in patients with COVID-19	
	Li J, Zhang YY, Cong XY, Ren SR, Tu XM, Wu JF	
	META-ANALYSIS	
585	Efficacy and safety of argatroban in treatment of acute ischemic stroke: A meta-analysis	
	Lv B, Guo FF, Lin JC, Jing F	



#### World Journal of Clinical Cases

#### Contents

Thrice Monthly Volume 10 Number 2 January 14, 2022

#### **SCIENTOMETRICS**

594 Biologic therapy for Crohn's disease over the last 3 decades Shen JL, Zhou Z, Cao JS, Zhang B, Hu JH, Li JY, Liu XM, Juengpanich S, Li MS, Feng X

#### **CASE REPORT**

- 607 Novel compound heterozygous GPR56 gene mutation in a twin with lissencephaly: A case report Lin WX, Chai YY, Huang TT, Zhang X, Zheng G, Zhang G, Peng F, Huang YJ
- 618 Patients with SERPINC1 rs2227589 polymorphism found to have multiple cerebral venous sinus thromboses despite a normal antithrombin level: A case report

Liao F, Zeng JL, Pan JG, Ma J, Zhang ZJ, Lin ZJ, Lin LF, Chen YS, Ma XT

Successful management of delirium with dexmedetomidine in a patient with haloperidol-induced 625 neuroleptic malignant syndrome: A case report

Yang CJ, Chiu CT, Yeh YC, Chao A

631 Malignant solitary fibrous tumor in the central nervous system treated with surgery, radiotherapy and anlotinib: A case report

Zhang DY, Su L, Wang YW

643 Anesthesia and perioperative management for giant adrenal Ewing's sarcoma with inferior vena cava and right atrium tumor thrombus: A case report

Wang JL, Xu CY, Geng CJ, Liu L, Zhang MZ, Wang H, Xiao RT, Liu L, Zhang G, Ni C, Guo XY

656 Full-endoscopic spine surgery treatment of lumbar foraminal stenosis after osteoporotic vertebral compression fractures: A case report

Zhao QL, Hou KP, Wu ZX, Xiao L, Xu HG

663 Ethambutol-induced optic neuropathy with rare bilateral asymmetry onset: A case report Sheng WY, Wu SQ, Su LY, Zhu LW

671 Vitrectomy with residual internal limiting membrane covering and autologous blood for a secondary macular hole: A case report

Ying HF, Wu SQ, Hu WP, Ni LY, Zhang ZL, Xu YG

677 Intervertebral bridging ossification after kyphoplasty in a Parkinson's patient with Kummell's disease: A case report

Li J, Liu Y, Peng L, Liu J, Cao ZD, He M

685 Synovial chondromatosis of the hip joint in a 6 year-old child: A case report Yi RB, Gong HL, Arthur DT, Wen J, Xiao S, Tang ZW, Xiang F, Wang KJ, Song ZQ

691 Orthodontic retreatment of an adult woman with mandibular backward positioning and temporomandibular joint disorder: A case report

Yu LY, Xia K, Sun WT, Huang XQ, Chi JY, Wang LJ, Zhao ZH, Liu J



Conter	World Journal of Clinical Cases ontents Thrice Monthly Volume 10 Number 2 January 14, 2022	
703	Autosomal recessive spinocerebellar ataxia type 4 with a <i>VPS13D</i> mutation: A case report <i>Huang X, Fan DS</i>	
709	Primary adrenal diffuse large B-cell lymphoma with normal adrenal cortex function: A case report <i>Fan ZN, Shi HJ, Xiong BB, Zhang JS, Wang HF, Wang JS</i>	
717	Varicella-zoster virus-associated meningitis, encephalitis, and myelitis with sporadic skin blisters: A case report	
	Takami K, Kenzaka T, Kumabe A, Fukuzawa M, Eto Y, Nakata S, Shinohara K, Endo K	
725	Tension pneumocephalus following endoscopic resection of a mediastinal thoracic spinal tumor: A case report	
	Chang CY, Hung CC, Liu JM, Chiu CD	
733	Accelerated Infliximab Induction for Severe Lower Gastrointestinal Bleeding in a Young Patient with Crohn's Disease: A Case Report	
	Zeng J, Shen F, Fan JG, Ge WS	
741	Occupational fibrotic hypersensitivity pneumonia in a halogen dishes manufacturer: A case report	
	Wang M, Fang HH, Jiang ZF, Ye W, Liu RY	
747	Using a fretsaw in treating chronic penial incarceration: A case report	
	Zhao Y, Xue XQ, Huang HF, Xie Y, Ji ZG, Fan XR	



#### Contents

Thrice Monthly Volume 10 Number 2 January 14, 2022

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CASE REPORT

# Occupational fibrotic hypersensitivity pneumonia in a halogen dishes manufacturer: A case report

Min Wang, Hao-Hui Fang, Zi-Feng Jiang, Wei Ye, Rong-Yu Liu

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### Abstract

#### BACKGROUND

Fibrotic hypersensitivity pneumonitis (FHP) is an allergic and diffuse pneumonia caused by repeated inhalation of antigenic substances, and sometimes developed in people working in specific environments. While novel antigens and exposures continued to be described, physicians should maintain a high suspicion of potential exposures. A detailed assessment of the patient's occupational exposures as well as living environment is necessary and complete allergen avoidance is the first and most important step in the management of FHP once the allergens are determined.

#### CASE SUMMARY

A 35-year-old female was admitted to the hospital with a cough and breathing difficulties for more than one year. She was a nonsmoker and a manufacturer of halogen dishes, which are characteristic Chinese foods, for 15 years without any protection. High resolution computed tomography of the chest demonstrated an interstitial pneumonia pattern. Pulmonary function examination showed restricted ventilation dysfunction and a significant reduction in dispersion ability. Cell differentiation in bronchoalveolar lavage fluid demonstrated lymphocytosis (70.4%) with an increased lymphocyte CD4/CD8 ratio (0.94). Transbronchial lung biopsy combined with lung puncture pathology showed diffuse uniform alveolar interval thickening, chronic inflammatory cell infiltration, a proliferation of tissue in the bronchial wall fiber and alveolar epithelial follicle degeneration, resulting in fibrosis.

#### **CONCLUSION**



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Exposure to spices used for the production of halogen dishes may cause FHP.

Key Words: Halogen dishes; Inhalation; Interstitial pneumonia; Spices; Fibrotic hypersensitivity pneumonitis; Case report

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Core Tip: Fibrotic hypersensitivity pneumonitis (FHP) is a disease related to immunological response to an inhaled antigen. Avoiding contact with susceptible antigen is the key. Here, we report a case of FHP in a halogen dishes manufacturer probably associated with exposure to the spices. We recommend to enhance respiratory protection during the production of such dishes to prevent this lung disease.

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#### INTRODUCTION

Fibrotic hypersensitivity pneumonitis (FHP) is a chronic and fibrosing interstitial lung disease caused by inhaled antigen exposure which can develop in people working in specific environments or with specific career[1]. The commonly known form is Farmer's lung[2]. Halogen dishes are Chinese cuisines in which long-term boiling of various spices powder is required before meat and vegetables are added. Each halogen maker has a so-called secret recipe, but almost all contain ginger, pepper, soy sauce, and other spices. A single case of hypersensitivity pneumonitis (HP) associated with curry powder and ground pepper in a potato chip factory has been described[3]. We herein report a case of FHP in a halogen dishes manufacturer, which may have been caused by the inhalation of spices.

FHP can lead to impaired respiratory function, reduced disease-related quality of life, and early mortality. Management of FHP should start with exposure remediation once the source of allergy is identified. A high index of suspicion is important to diagnosis, even after innocuous and apparently trivial exposures to potential antigens. We herein report a case of FHP in a halogen dishes manufacturer, which may have been caused by inhalation of spices.

#### **CASE PRESENTATION**

#### Imaging examinations

High resolution computed tomography (HRCT) of the chest showed thickening of interlobular septa and central lobular nodules which revealed honeycomb changes in severe areas (mainly in the lung base), as well as traction bronchiectasis (Figure 1).

#### Laboratory examinations

Routine blood, liver function, renal function tests and serum sodium, potassium, creatinine, magnesium, and calcium were all within normal limits. Blood tumor biomarkers including anti-nuclear antibody, anti-centromere antibody, anti-neutrophil cytoplasmic antibody, and Scl-70 antibody screens and C-reactive protein were negative. Pulmonary function tests showed a forced vital capacity (FVC) of 1.75 L (51.2% predicted), a forced expiratory volume in 1 s (FEV1) of 1.63 L (55%), an FEV1/FVC ratio of 109.6%, and carbon monoxide diffusing capacity 11.9 mmHg/mL/min (39% predicted). Cytology of bronchoalveolar lavage (BAL) fluid showed a large number of lymphocytes (the total number of cells was 4.16x10<sup>6</sup>/mL, lymphocytes accounted for 70.4%, of which the lymphocyte CD4/CD8 ratio was 0.94, and bronchoscopy was performed to rule out a superimposed infection. Allergen



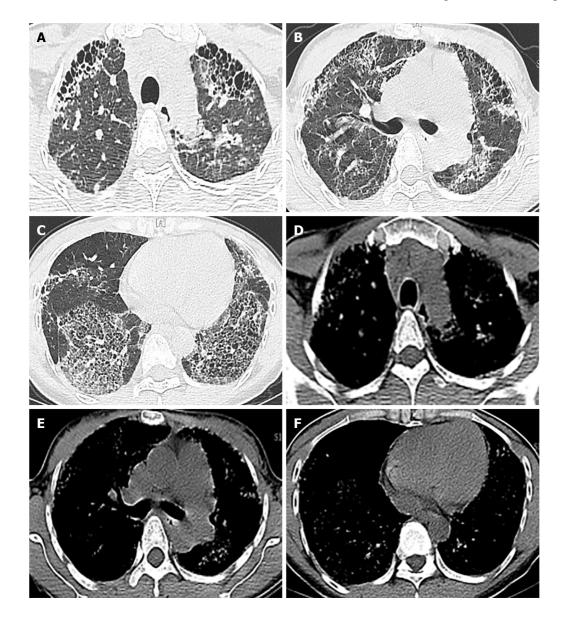


Figure 1 Chest high-resolution computed tomography (HRCT) images. A-C: showing traction bronchiectasis and extensive fibrosis, reticular changes (especially in the outer lung band and the double lower lung); D-F: showing the corresponding mediastinal window.

> determination showed that tree, pollen, ragweed, Artemisia argyi, home dust mite, home dust, cat hair, dog hair, cockroach, mold combination, legume, egg white, milk, peanut, soybean, beef, mutton, cod, shrimp, crab, salmon, lobster, scallop, etc. were all negative.

#### Physical examination

On examination, her temperature was 36.7 °C, blood pressure was 118/82 mmHg, heart rate was 74 bpm, oxygen saturation was 92% in room air, and respiratory rate was 22 breaths/min. The patient was identified as Grade 3-shortness of breath according to the mMRC (Modified Medical Research Council) dyspnea scale and lung examination revealed bilateral Velcro like crackles in the lower lung fields. The rest of her physical examination was unremarkable.

#### Personal and family history

This patient had no special personal and family history.

#### History of past illness

She was a self-employed worker, cooking and selling halogen dishes for over 15 years, without any measures to protect the respiratory tract. There was no history of orthopnea, chest pain, or paroxysmal nocturnal dyspnea. She denied smoking and there was no history of asthma, allergies, or any other prior pulmonary disease.



#### History of present illness

The patient took anti-inflammatory and cough medications intermittently, but there was no improvement in her symptoms.

#### Chief complaints

A 35-year-old Chinese woman with no significant medical history was admitted to our hospital because of a cough and breathing difficulties. She was a self-employed cook who produced and sold halogen dishes for more than 15 years without appropriate equipment to protect the airway from inhalation.

#### MULTIDISCIPLINARY EXPERT CONSULTATION

Fan-Qing Meng, MD, Professor, Department of Pathology. Fibrin-like exudation, more lymphocytes, foam-like tissue cells and a small number of macrophages were observed on histopathology of lavage fluid, suggesting persistent exposure to inhaled antigen. Transbronchial lung biopsy (TBLB) and computerized tomography-guided percutaneous lung puncture showed chronic lymphocyte and foamy macrophages infiltration, diffuse, relatively uniform alveolar septal thickness, hyperplasia of fibrous tissue in the bronchial wall, resulting in fibrosis (Figure 2).

#### FINAL DIAGNOSIS

Occupational FHP due to probable long-term exposure of spices in a halogen dishes manufacturer.

#### TREATMENT

The patient was started on prednisone 30 mg/d and pirfenidone anti-fibrosis treatment. Considering the side effects of high doses of hormones, the patient was discharged on a tapering schedule of prednisolone which was reduced by 5 mg every 2 wk.

#### OUTCOME AND FOLLOW-UP

The main symptoms of cough and breathing difficulties gradually improved slightly three months later, but no improvement in chest HRCT was noted. No significant adverse side-effects were noticed. The patient is currently on the lung transplant list.

#### DISCUSSION

This patient met the diagnostic criteria for FHP as she had a cough, breathing difficulties, long-term inhalation of spices, lymphocytic cellular pattern on BAL, bibasilar rales, reduction in diffusion capacity, and compatible radiologic and histopathological findings. BAL can not only exclude other major causes of interstitial pneumonia including alveolar hemorrhage, malignancy, infection, and eosinophilic lung disease, but also exclude common allergens. While BAL lymphocyte percentage in HP can vary substantially from 1.7% to 82% [4], in this patient BAL showed an obvious lymphocytic cellular pattern (70.4% lymphocytes) defined as a BAL differential count greater than 15%[5]. Additionally, aggregation of foamy macrophages and lymphocyte infiltration was observed which indicated exposure to inhalant pathogens [6]. Serologic studies for spices were not performed due to the patient's insurance status (she only had basic medical insurance and restricted economics), and had no relevance to the diagnosis even if negative as this would not exclude the clinical diagnosis of HP due to exposure to spices. Therefore, inhaled spices were the presumed cause of FHP in this case, especially given the absence of any other identifiable exposures.



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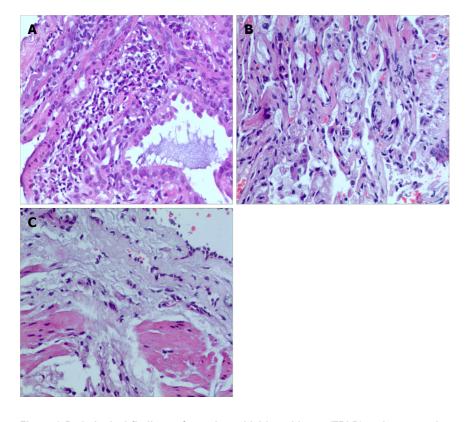


Figure 2 Pathological findings of transbronchial lung biopsy (TBLB) and computed tomography (CT)-guided percutaneous lung puncture, hematoxylin and eosin staining (×400). A: chronic lymphocyte infiltration; B: foamy macrophages infiltration and diffuse, relatively uniform alveolar septal thickness; C: hyperplasia of fibrous tissue in the bronchial wall, resulting in fibrosis.

Novel antigens and exposures related to occupation continue to be described, even innocuous and trivial exposures to potential antigens such as green tea[7], citrus[8], and pyrethrin<sup>[9]</sup> have been described. To our knowledge, this is the first case of FHP in a halogen dishes manufacturer caused by spices globally, which was confirmed by the presence of fibrosis through histological evaluation of TBLB and percutaneous lung puncture. However, it has also been pointed out that in the presence of a concomitant allergic predisposition, such as bronchial asthma, allergic rhinitis, or chronic bronchitis, the biological response to inhalation may be enhanced. There was no apparent comorbidity in our patient and she was a nonsmoker.

Heightened awareness of potential exposures is imperative in making a diagnosis and a detailed assessment of occupational exposures as well as the occupational environment is necessary as demonstrated in this case. Avoiding contact is the key for patients with HP, not only for patients but also for other individuals who may be exposed to the same allergen[9]. Although there are no obvious symptoms or failure of regular occupational disease physical examination or disease detection, it is essential to remind workers of their safety during the production process<sup>[8]</sup> (respirator, masks, improve ventilation, regular monitoring of air quality) and the compliance of workers should be confirmed. For those with established allergies, there are various ways to avoid exposure, for example, by changing jobs or occupations. Clinically appreciable improvement in symptomatic, physiologic, and radiographic features may be seen only in patients with non-fibrotic HP[10]. Due to the long and insidious exposure to antigen, fibrosis was observed both radiologically and pathologically in our patient. Management of FHP should start with exposure remediation where possible, oral glucocoticoids and azathioprine achieved an initial improvement in pulmonary function tests and symptoms[11], anti-fibrosis therapy should be considered in patients with symptomatic or progressive disease, and non-pharmacologic and supportive management should be offered and, in cases of treatment-resistant, progressive illness, lung transplant should be considered[12,13]. Based on this, we advised our patient to change her occupation, and prednisone and anti-fibrosis therapy was given to improve her symptoms. Halogen dishes manufacture is mostly a family business in China, the patient inherited the shop from her mother and had no other life skills, although she was no longer responsible for food preparation, she still sold the halogen dishes at the counter. Following treatment, her symptoms improved slightly and requiring long-term oxygen therapy, the HRCT scan showed no



improvement. The patient is currently on the waiting list for a lung transplant.

#### CONCLUSION

Attention should be paid to the possibility of inhalation hypersensitivity pneumonitis caused by spices. Improvements in respiratory protection measures should be suggested for workers during the production process to avoid this type of lung disease.

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