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Chronic Pulmonary Aspregillosis (CPA)







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19/7/2013: Admission to the hospital :

- Haemoptysis
- Cavity lession
- Checked for TB
- (+) Mantoux
- Positive galactomannan antigen bronchial (BAL)/serum
- CT angiography. Absence of visible bleeding site
- Voriconazole

18/10/2013 : New bleeding

18/03/2014: Discontinuous treatment for Aspergilossis





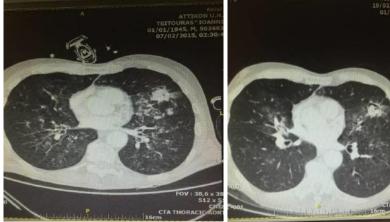


Male 69 yrs

28/01/2015 : 3 episodes of massive haemoptysis **06.02.2015** : Admission to ICU due to:

- o massive haemoptysis
- o aspiration
- o desaturation
- o loss of consciousness
- o Intubation- mechanical ventilation

Ο



- o 2006: prostate cancer, prostatectomy
- o 2013: radiotherapy 03-23/05/2013
 - 2013: nephrotic syndrome / AKF
 - kidney biopsy → diffuse proliferative glomerulonephritis
 - IGC: immunocomplex associated immunodifficiency /chronic thrombotic microangiopathy possibly due to radiotherapy
- Methylprednisolone and cyclophosphamide.
- o Oral methylprednisolone



During the next two years

Symptoms

- o Weakness
- o Weight loss
- o Haemoptysis

<u>Tests</u>

- 8 CT and CTA
- Never embolized

Therapeutic interventions in summary

- o Voriconazole 07/13-03/14 (9 months)
- o Patient stops therapy for 4 months
- Restarted voriconazole 07/14 stops in 10/14 (3 months)
- Patient stops therapy again for 2 months
- Haemoptysis 04/01/15 voriconazole was restarted
 - New episode of haemoptysis
 - Admission to the Internal Medicine Clinic



- o Admission to ICU 06/02/2015
- o Sedation
- o Analgesia
- o Intubation Ensuring airway

Patient problems

- * Frequently hemoptysis
- * Imminent airway
- Inadequate ventilation and oxygenation

Actions performed

- o Double lumen intubation tube
- o Bronchoscopy
- Early tracheostomy day 5

Diagnostic tests performed

- TB / Nocardia (-)
- Biopsy nasal mucosa (-)
- GM serum and bronchial aspirates (+)
- Precipitins : IgG 6.5 / IgM 0.4 / IgA 1.1
- Biopsy bronchial mucosa (-)
- Lung Biopsy (Confers aspergillosis)
- Voriconazole





06/03 Surgical removal of the cavity Patient returned to the ICU

16/03 surgical stump opens bronchopulmonary fistula

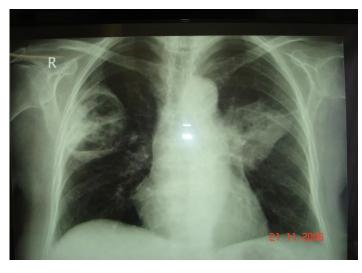
27/04 surgical closure of bronchopulmonary fistula with pericardial patch



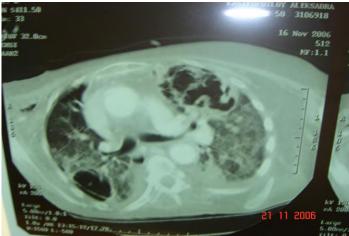
The patient after 4 months of hospitalization in the ICU died due to septic shock and multiorgan failure (MOF)



Chronic Pulmonary Aspergillosis (CPA) CPA- Imaging findings / SAIA







Female patient with leukemia ARDS / Resp failure Intubation Mechanical Ventilation BAL : Aspergillus / GM (+) Voriconazole Death Autopsy : SAIA



Chronic Pulmonary Aspergillosis (CPA) CPA- Treatment

- 1. Treatment with Antifungals
- 2. Surgical treatment
- 3. Treatment of hemoptysis

a. Local cavity therapy

- 4. Corticosteroids
- 5. INF-γ



Chronic Pulmonary Aspergillosis (CPA) CPA – Treatment with antifungals

PO triazole therapy

- Patient's type of disease or clinical phenotype and eligibility for surgical treatment

Main target : QUALITY OF LIFE

- CCPA : Voriconazole or itraconazole PO Duration : at least 4-6 months, slow response
- CFPA : Long-term itraconazole PO → stabilizing patients
- **SAIA** : Treatment as in IPA

IV administration

WHERE ?

Progressive disease, PO failure, intolerant of triazoles, resistance

WITH WHAT ?

IV **Micafungin** 150mg/day vs **Voriconazole** (response to treatment vs 60 vs 53%), **Liposomal AmB** 3 mg/kg/ day, **Caspofungin** 50–70 mg/day



Chronic Pulmonary Aspergillosis (CPA) CPA – Surgical treatment

- 1. Aspergilloma : in patients with adequate pulmonary function
 - a. Full resection without spillage of fungal elements into the pleural space.
 - b. Recurrence of disease and haemoptysis are rare in simple aspergilloma
- 2. CCPA carries a lower success rate
- 3. Surgery : in all patients with severe haemoptysis
- 4. Careful patient selection to avoid peri- and post-operative complications
 - a. Bullectomy, segmentectomy, sublobar resection, wedge resection, lobectomy, pleurectomy, pneumonectomy

5. Post-operative complications

a. persistent air-leak, empyema, pneumonia, wound infection, bronchopleural fistula, respiratory failure, massive haemorrhage, and death

6. Catheter embolisation of bronchial arteries

a. prior to surgery and a bridge towards selective surgery



Chronic Pulmonary Aspergillosis (CPA) CPA – Local cavity therapy

Aim : to control recurrent haemoptysis if surgical treatment is not an option in those without a haemorrhagic diathesis when systemic use of antifungals is ineffective or prevented by Aes

- Instillation of antifungals through

- an endobronchial catheter under bronchoscopic guidance
- via a percutaneous transthoracic needle or
- catheter placed into the aspergilloma cavity

- Antifungals

- Amphotericin B (as paste or solution), the drug of choice (50 mg in 20 mL 5% Dextrose solution)
- azoles (miconazole, itraconazole)
- sodium iodide and nystatin (as paste with amphotericin B).

- Complications

- cough, chest pain, pneumothorax or endobronchial reflux
- Short-term response rates : 70% to 100%.



Chronic Pulmonary Aspergillosis (CPA) CPA – Therapies for haemoptysis

I. Mild, moderate or life-threatening haemoptysis

- CCPA and simple aspergilloma usually
- Tranexamic acid (typically 500 mg three times daily)

II. Moderate or severe haemoptysis

- Embolisation either as a temporising measure before surgery or as a definitive treatment

III. Complications of embolisation

- chest wall pain, stroke with cortical blindness or impaired vision
- chest wall or spinal cord infarction, as well as renal impairment and allergic reactions to the contrast dye.



Chronic Pulmonary Aspergillosis (CPA) CPA – Corticosteroids / IFN-γ

Patients with underlying diseases

- sarcoidosis, rheumatoid arthritis, COPD, ABPA or asthma may be dependent on corticosteroids
- Prednisolone 5–30 mg/day may carefully be considered for symptom control only if are adequately treated with antifungals

Interferon (IFN)-γ deficiency

- Impaired production of IFN-γ and IL-12 necessary to produce IFN-γ
- IFN-γ substitution (Adjunctive therapy) with 50–60 µg subcutaneously, three times weekly
- Clinical improvement



Chronic Pulmonary Aspergillosis (CPA) CPA – Follow up

After resection surgery of Aspergillus nodule

1. Single and completely excised aspergilloma

The patient does not require antifungal therapy unless immunocompromised

2. A single nodule not completely resected

Quantitative Aspergillus IgG serology, inflammatory markers and radiology at 3-monthly intervals to determine if antifungal therapy required

3. Multiple nodules

Antifungals • reduction in size of most or all nodules over time An increase in size may represent another disease process, such as a malignancy

4. Close radiological follow-up (initially 3 monthly)

To ensure there has been no progression

5. In all cases, corticosteroid exposure should be minimised