A strange case of deep venous thrombosis (DVT)

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Brasilian woman, 50 years old In Italy for 10 years (last trip to Brazil 6 months before) Pain and swelling of the left lower limb

Medical history:

- 4-year history of post-prandial abdominal pain, constipation and 7 kg weight loss (reduced food intake) → upper and lower endoscopy and barium enema negative
- Left ovaric cyst in follow-up
- Kidney stones
- Menopause state, no HRT

- No therapy
 - No allergy
- No smoking

On Admission

- ✓ BP 80/40 mmHg, HR 80 b.p.m rt, SatO2 97% AA; TA 36° C, RR 16/min; BMI 18 kg/m2.
- ✓ Normal ECG
- ✓ Physical examination: mild diffuse abdominal discomfort and tenderness, especially in left iliac fossa; edema of the lower left limb
- ✓ Neurological examination: normal

Other investigations:

- ✓ Chest X-ray: nodule in the upper right lung (result of past infection?)
- ✓ Doppler US lower limbs: <u>Left deep venous</u>
 <u>femoro-popliteal thombosis</u> (DVT) →
 treatment with subcutaneuos nadroparin and
 compression stockings

Bloods	
Hb (g/dl)	10,5
MCV (fl)	84
WC (mmc)	13250
PTL (mmc)	350000
Creatinine (mg/dl)	0,6
AST (U/L)	20
ALT (U/L)	25
GGT (U/L)	15
Albumin	3,2
Cholinesterase (U/L)	3364
Total cholesterol (mg/dl)	110
CRP (mg/dl)	6,6
Procalcitonin (mg/dl)	0,08
PT	1,24
PTT	0,92
D-dimer (μg/ml)	923
LDH (U/L)	110

Causes of DVT

Risk factors (causes) for the development of venous thrombosis

Inherited thrombophilia
Factor V Leiden mutation
Prothrombin gene mutation
Protein S deficiency
Protein C deficiency
Antithrombin (AT) deficiency
Rare disorders
Dysfibrinogenemia
Acquired disorders
Malignancy
Presence of a central venous catheter
Surgery, especially orthopedic
Trauma
Pregnancy
Oral contraceptives
Hormone replacement therapy
Tamoxifen, Thalidomide, Lenalidomide
Immobilization
Congestive failure
Antiphospholipid antibody syndrome
Myeloproliferative disorders
Polycythemia vera
Essential thrombocythemia
Paroxysmal nocturnal hemoglobinuria
Inflammatory bowel disease
Nephrotic syndrome

Also:

- Obesity
- Smoking
- Chronic inflammatory state

But in this case we excluded:

- Smoking
- Obesity
- Hormonal therapy
- History of surgery, trauma or immobilization
- Familiarity for DVT or abortion
- IBD (endoscopy)

Which exams would you ask for?

✓ Inflammatory markers: Blood count (WC 8000-12000/mmc), CRP 3-6 mg/dl, Procalcitonin 0,04-0,08 mg/dl → stable

✓ Thrombophilic screening: ANA, Ab anti DNAds, protein C and protein S levels, Antitrombin level, Antiphospholipid antibodies (LLAc, ACA IgM, Ab anti beta₂-microglobulin), Factor V Leiden mutation, prothrombin gene mutation, homocystein level → negative

✓ Neoplastic markers: CEA, Ca 19.9, AFP, Ca 125, Ca 15.3 → negative

During the following day in the ward

Persistent and worsened abdominal pain, BP 90/60 mmHg, HR 90 rt, SpO2 94% AA, TA 36° C. Mild dyspnea. Normal ECG and BGA.

What do you suspect (DVT)? What do you do???

* Thorax / coxofemoral CT: exclusion of pulmonary embolism (PE) BUT evidence of two solid hypodense lesions in the upper and inferior lobe of the right lung characterized by contrast enhancement. Large hypodense lesion in the ileopsoas muscle bilaterally. Similar lesions in the left ovarian and obturator area, in the Pouch of Douglas and at the hepatic hilum (neoplastic vs infectious?).

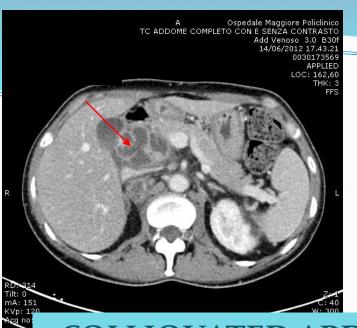
AND NOW?

❖ <u>Abdomen CT</u>: complex hypodense septed cystoid lesion at the hepatic hilum, between head of pancreas, portal vein and IVC, similarly large hypodense lesions in the ileopsoas muscle bilaterally (left 3x 2,5x8 cm), along the left common iliac artery and left external iliac artery (diam max 4 cm) and in the left ovarian area. Similar lesion in the inferior lobe of the right lung. (NEOPLASTIC vs INFECTIOUS)

How to better characterize these complex lesions?

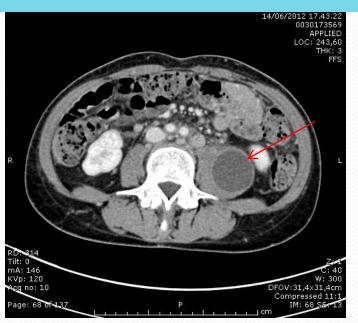
CT/PET total body

actively inflammatory nodal colliquated lesions (SUV max 6,3)





COLLIQUATED ABDOMINAL LYMPHADENOPATHIES AND BILATERAL PSOAS ABSCESS





Lymphadenopathy

- ✓ Infectious disease:
 - Bacteria, virus, mycobacteria, fungi, protozoa, spirochaete, parasites
- ✓ Neoplastic disease
 - Head/neck squamocellular carcinoma
 - Metastasis
 - ✓ Lymphoproliferative disease
 - ✓ Immunological disease (serum sickness, drug reaction)
 - ✓ Miscellany (sarcoidosis, SLE, RA, Still's disease, Dermatomyositis, Churg-Strauss syndrome, Castelman syndrome)

Psoas Abscess

- ✓ Infectious disease
 - Primary (20%): S.Aureus (80%), immunocompromised (dialisis, CT,NPL, AIDS, DM)
 - Secondary (80%): direct diffusion from a focus near by → especially G.I in Chron disease; E.coli, M.TBC, Streptococcous
- ✓ Neoplastic disease
 - Direct diffusion from retroperitoneal tumors (scheletric or colic/urogenital)
 - Primary (lipo/fibrosarcoma)
 - Metastasis

CT appearence similar to abcess

- ✓ Others
 - Hemorragic lesions (oral anticoagulation, spontaneous rupture of vascularised neoplastic lesion or aneurisma; trauma)
 - Iatrogenic post urologic or orthopaedic surgery

What's next?

- 1. Serologic screening for infection
- 2. Autoimmune screening
- 3. Neoplastic and haematologic markers
- 4. Quantiferon-TC Gold In-tube assay
- 5. Drainage of the lesions
- 6. All them

Serological investigations

- ✓ Serology for
 - •Virus: HIV,HCV,HBV neg
 - •Fungal, protozoa, parasites:
 - Toxoplasma gondii neg
 - Trypanosoma cruzii neg
 - Cysticercosis neg
 - Trichinella spiralis neg
 - Echinococcus (IgG pos, Ig M doubt)
 - Histoplasma capsulatum neg
 - •Mycobacterium tbc: Quantiferon-TC Gold In-tube assay positive
- ✓ Markers npl: CEA, Ca19.9, Ca125, Ca15.3 neg
- ✓ Autoimmune screening: ANA 1:80 (speckled granular)
- ✓ LDH and protein electrophoresis neg b2microglobuline 3.9 µg/ml

Previous tubercular contact

Doubt echinococcal serology

Which is your diagnosis?

- 1. Piogenic abscesses (S. aureus, E.coli..)
- 2. Tubercular infection
- 3. Echinococcal infection
- 4. Solid neoplastic disease
- 5. Lymphoproliferative disease
- 6. Sarcoidosis

WOULD YOU DRAIN THE LESION???

Echo-guided percutaneous fine needle aspiration of the psoas abscess: <u>PUS COLLECTION</u>

- → Coltures negative for bacteria, myceti or parasites
- → Microscopy examination with Ziehl-Nielsen stain negative for Mycobacterium tbc (acid fast bacilli)
- → Colture and PCR for Mycobacterium tbc in progress(Quantiferon positive)
- →Cytopathological examination not diagnostic (purulent sample)

AND NOW????

Echoendoscopy of the lesion between hepatic hilum and pancreas: lymph nodes with max diam 8 cm and necrotic aspect → aspiration of nodal material BUT INADEQUATE SPECIMEN.

In the meanwhile..

- Patients still symptomatic for abdominal pain, responsive to paracetamol
- Partial resolution of the left femoro-popliteal thrombosis (US Doppler) but disappearence of the edema of the left lower limb
- Still on anticoagulant therapy (nadroparin 0,5 twice a day)

Colture of the psoas specimen positive for Mycobacterium tuberculosis

→ patient referred to an antitubercular Centre (Villa Marelli, H. Niguarda, Milan) and started to a quadruple therapy with ethambutol 1200 mg, rifampin 600 mg, isoniazid 250 mg and pyrazinamide 1500 mg

DIAGNOSIS: ABDOMINAL NODAL TUBERCULOSIS AND BILATERL TUBERCULAR ILEOPSOAS ABSCESS

HOW SYMPTOMS COULD BE EXPLAINED?

Abdominal lymph nodes involvement

Bowel compression by lymphoadenopathies

Long history of abdominal pain, especially post prandial episodes, associated with constipation and weight loss

Haematogenous spread or via direct proximity from iliac lymph nodes to the ileopsoas muscular tissue

Compression by ileopsoas abscess

Proinflammatory and procoagulatory state caused by infection

Femoro-popliteal venous thrombosis

Virchow-'s triad

- . **Direct endotelial injury** by the mycobacterium
- 2. Hemodynamic changes: stasis cause of external compression by lymph nodes and muscular abscesses
- 3. **Hypercoagulability** (↓ PC, PS, ATIII and ↑ PAI-1, fibrinogen and FVIII; Anticardiolipin antibodies; Mycobacterium tbc induces macrophages to produce pro-aggregating cytokines (IL6, IL1, TNF).

Incidence of DVT associated with TBC is nearly 1% (30% PE)



Thank you for your attention!!!!

