



Université  
Paris Cité



## Infections abdominales et immunodépression

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# Causes de l'immunodéficience

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- ▶ Génétique
- ▶ Physiologie
- ▶ Acquis
- ▶ Maladies chroniques
- ▶ Traitement (iatrogène)
- ▶ Hématologie



Infections du tractus digestif

Infection des organes solides

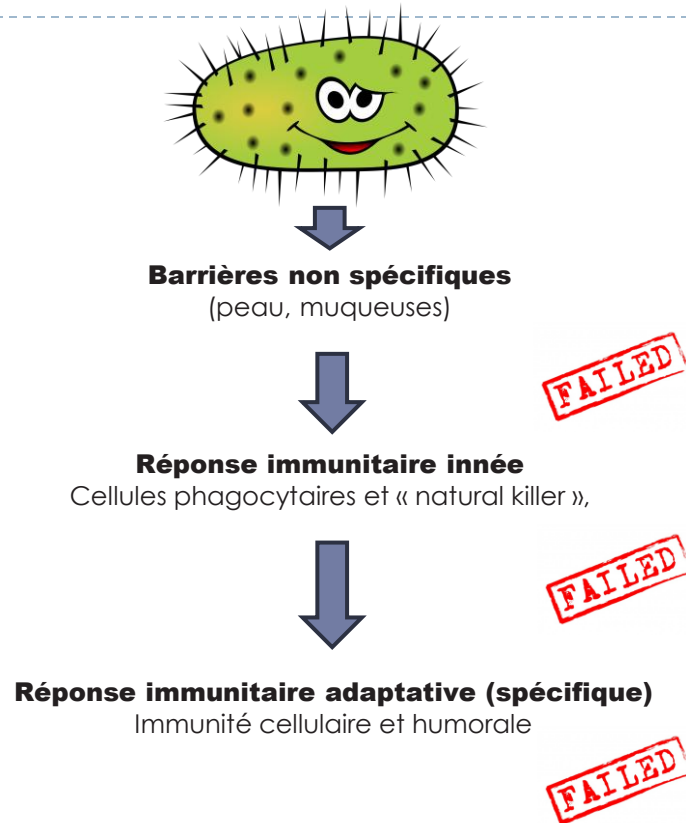


# Infections du tractus digestif

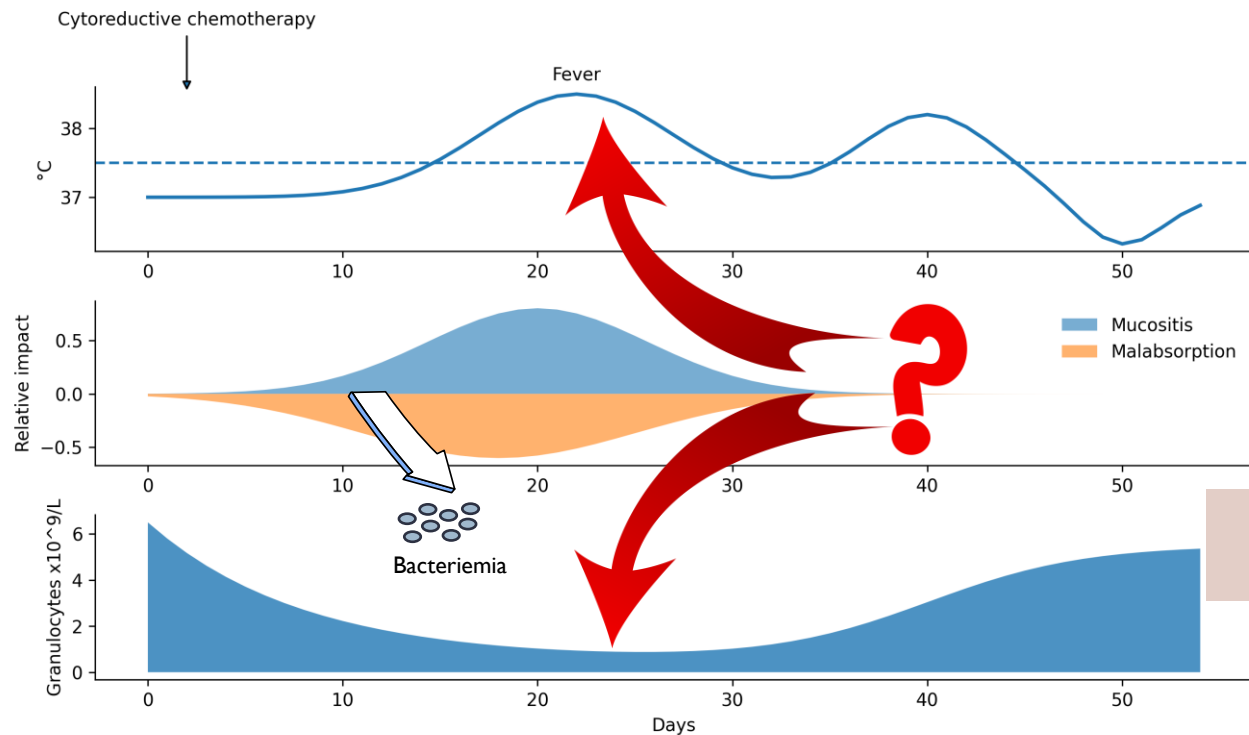
Principalement des colites

# Système immunitaire et défense contre les pathogènes

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# Chimiothérapie



C'est ici qu'arrive la  
demande de scanner ...

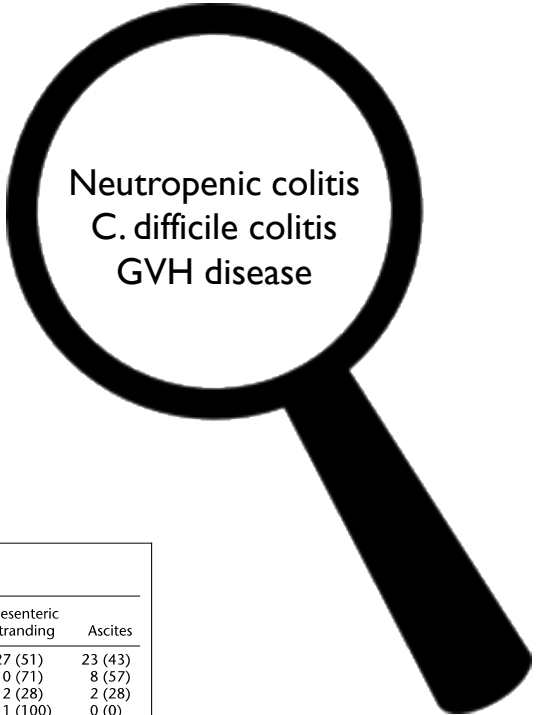
Iain D. C. Kirkpatrick, BSc,  
BSc(Med), MD  
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**Index terms:**

Colitis, 75.2043, 75.206  
Colitis, pseudomembranous, 75.263  
Enteritis, 73.69, 74.69  
Intestines, ischemia, 73.266, 74.266

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Radiology 2003; 226:668–674

# Gastrointestinal Complications in the Neutropenic Patient: Characterization and Differentiation with Abdominal CT<sup>1</sup>



Neutropenic colitis  
C. difficile colitis  
GVH disease

- ▶ 76 neutropenic patients with radiologic bowel abnormalities

**TABLE 2**  
Number of Patients with CT Abnormalities of Bowel Wall

| Diagnosis                                        | No. of Patients | Wall Thickening | Wall Nodularity | Mucosal Enhancement* | Pneumatosis |
|--------------------------------------------------|-----------------|-----------------|-----------------|----------------------|-------------|
| Neutropenic enterocolitis                        | 53              | 53 (100)        | 1 (2)           | 10 of 35 (28)        | 11 (21)     |
| <i>C difficile</i> colitis                       | 14              | 14 (100)        | 5 (36)          | 2 of 11 (18)         | 0 (0)       |
| Acute gastrointestinal graft-versus-host disease | 7               | 6 (86)          | 0 (0)           | 5 of 7 (71)          | 0 (0)       |
| Cytomegaloviral colitis                          | 1               | 1 (100)         | 1 (100)         | 0 of 0 (0)           | 0 (0)       |
| Bowel ischemia                                   | 1               | 1 (100)         | 0 (0)           | 0 of 0 (0)           | 1 (100)     |

Note.—Numbers in parentheses are percentages of diagnostic group unless otherwise noted.

\* Percentages were calculated by using only the number of patients who received intravenous contrast material.

**TABLE 3**  
Number of Patients with CT Abnormalities Not of the Bowel Wall

| Diagnosis                                        | No. of Patients | Bowel Dilatation | Mesenteric Stranding | Ascites |
|--------------------------------------------------|-----------------|------------------|----------------------|---------|
| Neutropenic enterocolitis                        | 53              | 20 (38)          | 27 (51)              | 23 (43) |
| <i>C difficile</i> colitis                       | 14              | 2 (14)           | 10 (71)              | 8 (57)  |
| Acute gastrointestinal graft-versus-host disease | 7               | 6 (86)           | 2 (28)               | 2 (28)  |
| Cytomegaloviral colitis                          | 1               | 0 (0)            | 1 (100)              | 0 (0)   |
| Bowel ischemia                                   | 1               | 1 (100)          | 0 (0)                | 1 (100) |

Note.—Numbers in parentheses are percentages of diagnostic group.

# Neutropenic Enterocolitis: An Uncommon, but Fearsome Complication of Leukemia

Rodrick Babakhanlou<sup>a, c</sup>, Farhad Ravandi-Kashani<sup>a</sup>,  
Dimitrios P. Kontoyiannis<sup>b</sup>

**Table 1.** Differential Diagnosis of Abdominal Pain in Neutropenic Patients [26]

CMV colitis

*C. difficile* colitis

Leukemic infiltrates of the bowel

Graft-versus-host disease

Ischemic colitis

Cholangitis

Cholecystitis

Acute appendicitis

Bowel obstruction

Autoimmune colitis due to check point inhibitors



# Colite neutropénique

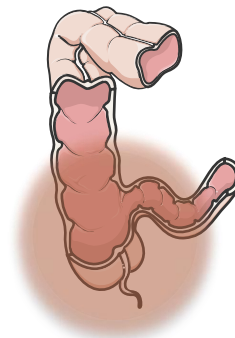
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## ► Synonymes

- Entérocolite nécrosante
- Typhlite = Syndrome iléo-caecal

## ► Contexte

- 5% des patients atteints d'hémopathies malignes, également patients traités par des chimiothérapies à haute dose pour des tumeurs solides
- Il faut l'association de la neutropénie (hemopathies +++) ET d'une altération de l'intégrité muqueuse due à la chimiothérapie cytotoxique



# Colite neutropénique

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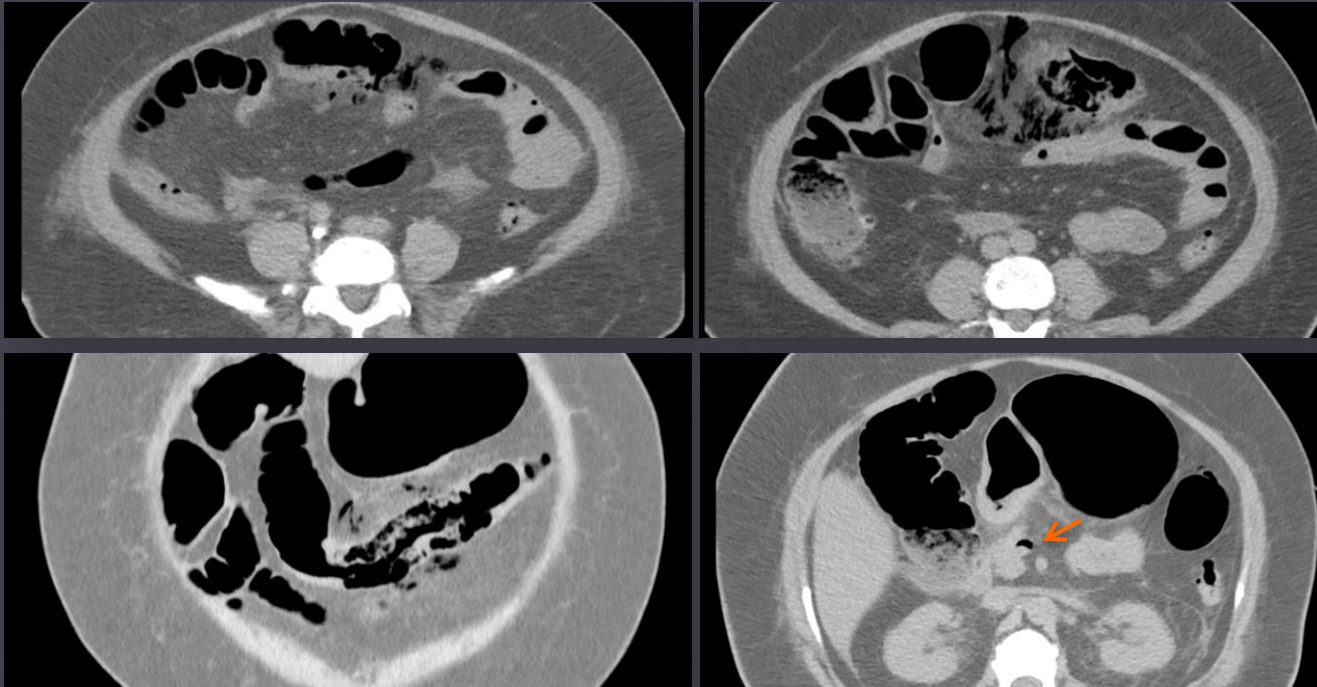
- ▶ Epaissement pariétal
- ▶ Rehaussement muqueux
- ▶ Infiltration mésentérique
- ▶ Pneumatose

Typhlite



## Colite neutropénique

Aplasie fébrile et neutropénie. Diabète. Pneumatose avec perforation. Présence d'air dans la VMS.

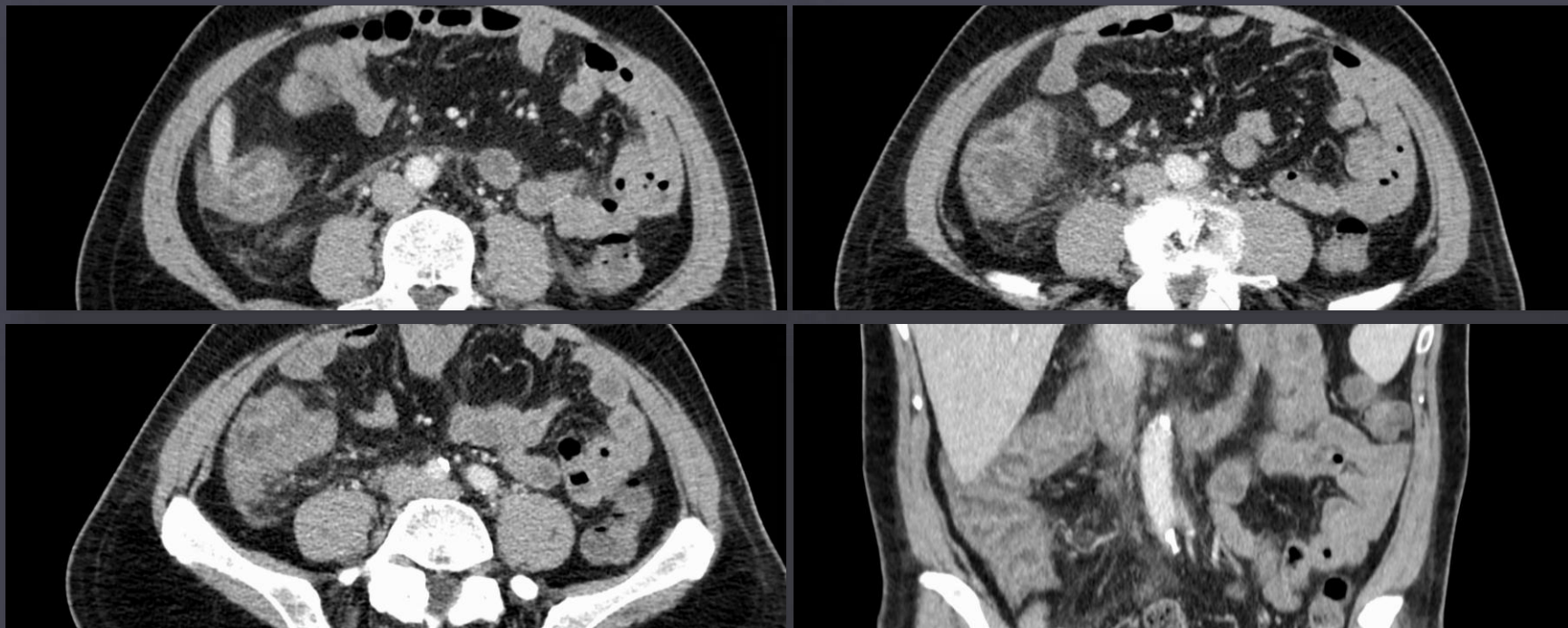


## Colite neutropénique

Patient transplanté rénal. PTLD en début de traitement. Colite perforée.



LAM, 3 semaines post induction. Neutropénie fébrile et diarrhée



# Colite pseudo-membraneuse

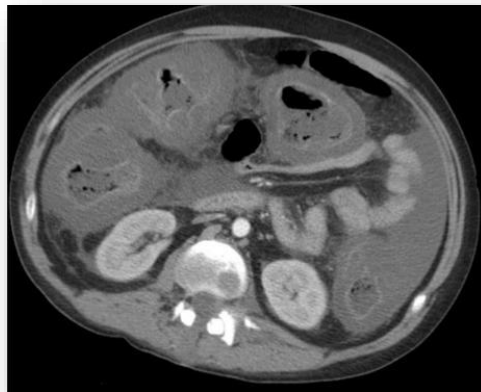
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- ▶ Colite infectieuse aiguë causée par des toxines produites par *Clostridium difficile*
- ▶ Complication Classique des traitements antibiotiques
- ▶ Diarrhée, douleurs abdominales, fièvre, hyperleucocytose

Complication de l'ABthérapie de  
prévention de la colite neutropénique !

## ▶ CT

- ▶ Epaissement pariétal majeur avec rehaussement muqueux
- ▶ 'Target sign' ou 'double halo sign', 'accordion sign'
- ▶ Peu d'infiltration de la graisse, ascite

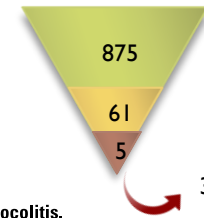


# Clostridium difficile Infection in Patients with Neutropenia

Marcus Gorschlüter,<sup>1</sup> Axel Glasmacher,<sup>1</sup> Corinna Hahn,<sup>1</sup> Frank Schakowski,<sup>1</sup> Carsten Ziske,<sup>1</sup> Ernst Molitor,<sup>2</sup> Günter Marklein,<sup>2</sup> Tilman Sauerbruch,<sup>1</sup> and Ingo G. H. Schmidt-Wolf<sup>1</sup>

Departments of <sup>1</sup>Internal Medicine I and <sup>2</sup>Medical Microbiology and Immunology, University of Bonn, Bonn, Germany

- ▶ 875 patients receiving cycles of myelosuppressive therapy
- ▶ 61 independent episodes of *Clostridium difficile*-associated diarrhea
- ▶ 5 developped severe enterocolitis

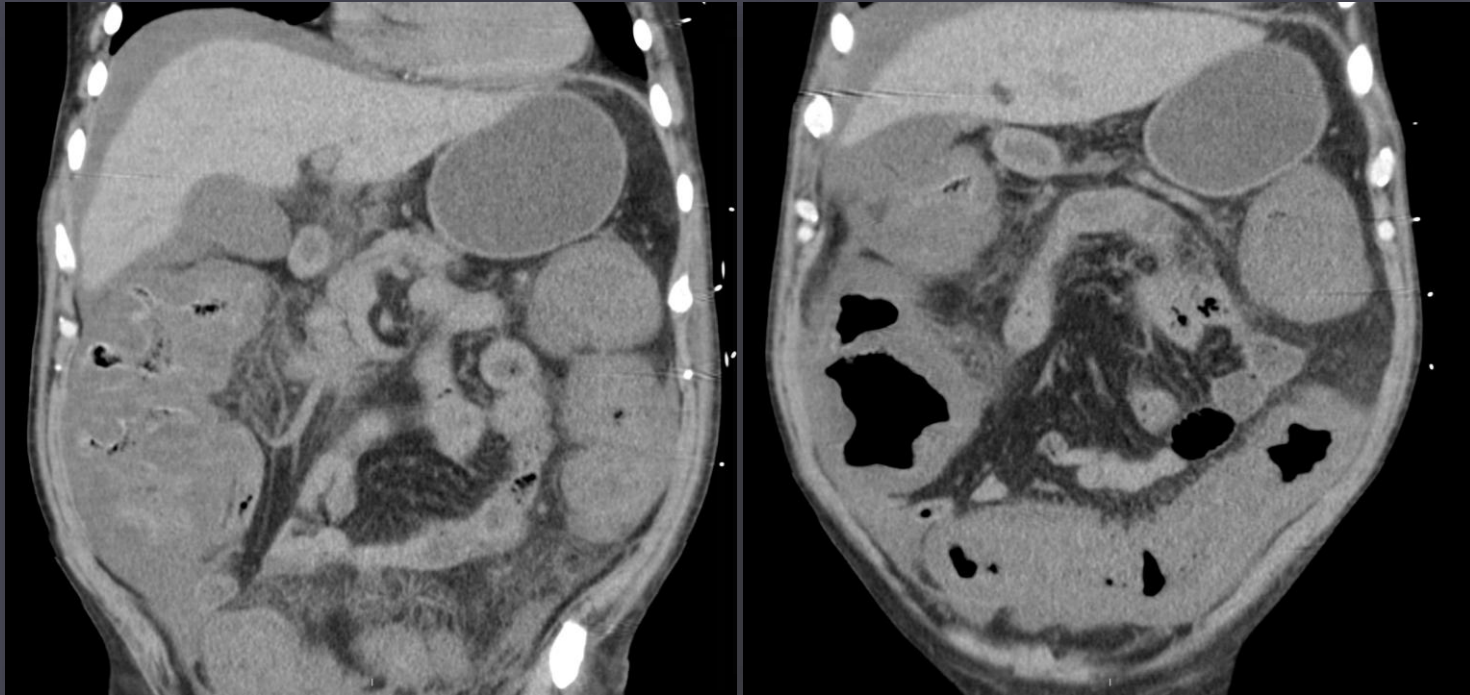


**Table 4. Characteristics of patients with *Clostridium difficile*-associated diarrhea (CDAD) who developed severe enterocolitis.**

| Patient no. | Sex, age in years | Underlying disease                      | Primary pathogenic cause of severe enterocolitis | Clinical symptoms                                  | Outcome/cause of death           | Therapy for CDAD                           |
|-------------|-------------------|-----------------------------------------|--------------------------------------------------|----------------------------------------------------|----------------------------------|--------------------------------------------|
| 11          | F, 62             | Chronic myeloid leukemia (blast crisis) | <i>Candida glabrata</i>                          | Fever, diarrhea, nausea, vomiting                  | Death/septic shock and pneumonia | iv metronidazole                           |
| 16          | M, 27             | Acute myeloid leukemia                  | Cytomegalovirus                                  | Fever, diarrhea, nausea                            | Recovery                         | Oral and iv metronidazole, oral vancomycin |
| 17          | M, 59             | Acute lymphoblastic leukemia            | <i>Candida krusei</i>                            | Fever, diarrhea, nausea, abdominal pain            | Death/septic shock               | iv metronidazole, oral vancomycin          |
| 23          | F, 39             | High-grade non-Hodgkin's lymphoma       | <i>C. difficile</i>                              | Fever, diarrhea, nausea, meteorism, abdominal pain | Death/pneumonia                  | iv metronidazole, oral vancomycin          |
| 44          | F, 18             | Acute myeloid leukemia                  | <i>C. difficile</i>                              | Fever, diarrhea, nausea, vomiting, abdominal pain  | Recovery                         | iv metronidazole                           |

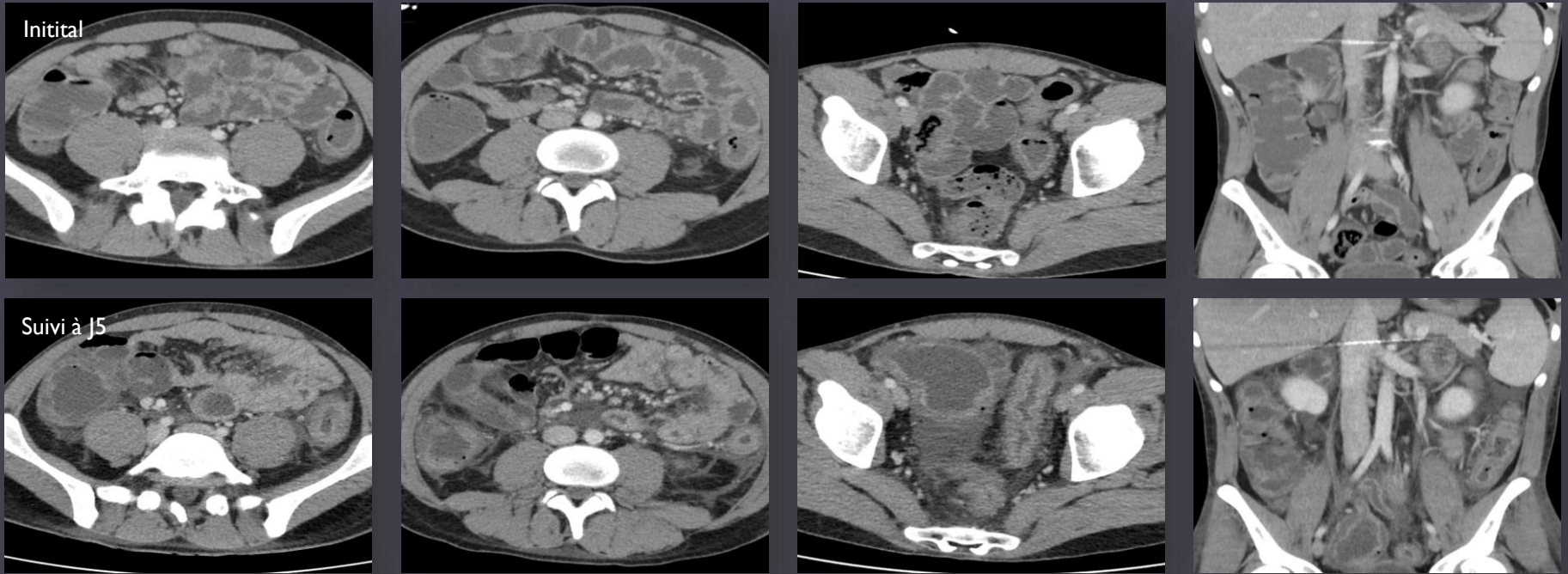
## Colite à Clostridium difficile

Myélome multiple en cours de chimiothérapie. ABthérapie à large spectre. Douleurs abdominales,,fièvre. 'Accordion sign', ascite.



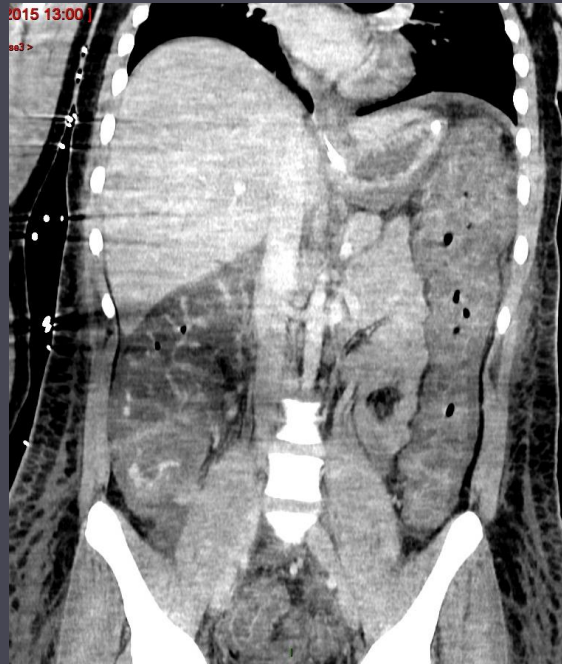
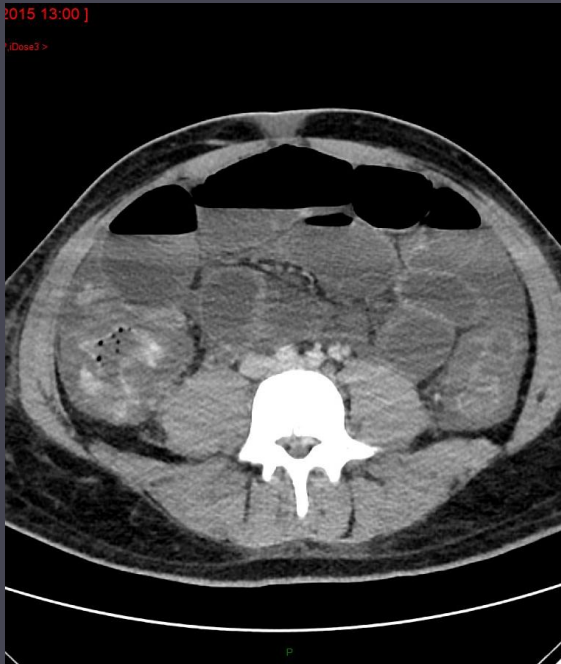
## Colite à Clostridium difficile

DLBCL traité par R-DHAX. Neutropénie fébrile traitée par AB. Diarrhée à Clostridium difficile suivie d'une colite sévère



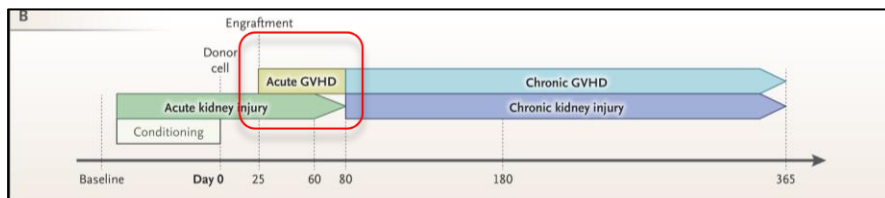
## Colite à *Clostridium difficile*

LAM 4. Aplasie fébrile depuis J20 post-induction. Antibiothérapie prolongée. Diarrhée à J40.



# Acute Graft versus Host Disease (GvHD)

- ▶ Surtout greffe de moelle
- ▶ Mismatch donneur/receveur HLA
- ▶ Entre J25-J80 après la greffe
- ▶ Diarrhée aqueuse > 500ml
- ▶ Douleurs abdominales sévères
- ▶ Iléus et diarrhée sanglante



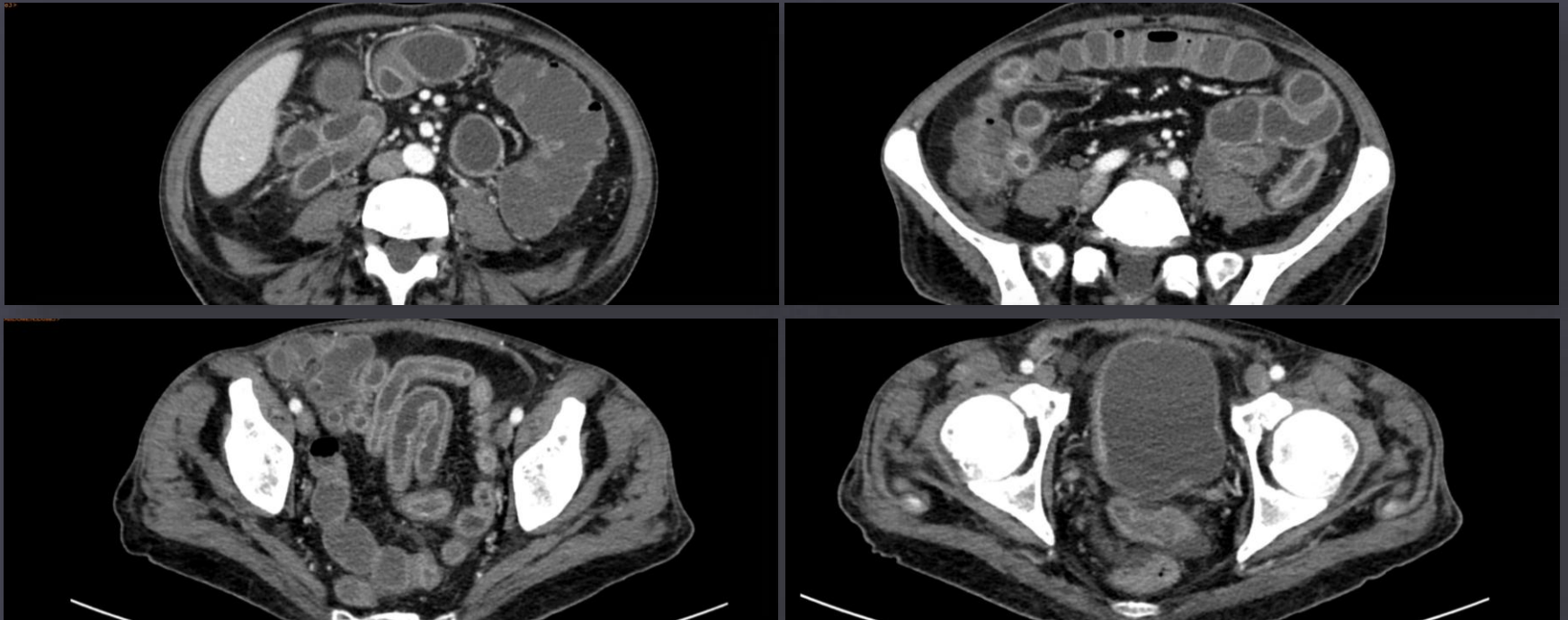
Babak N. Kalantari<sup>1</sup>  
 Koenraad J. Mortelé<sup>1</sup>  
 Vito Cantisani<sup>1</sup>  
 Silvia Ondategui<sup>1</sup>  
 Jonathan N. Glickman<sup>2</sup>  
 Adheet Gogate<sup>1</sup>  
 Pablo R. Ros<sup>1</sup>  
 Stuart G. Silverman<sup>1</sup>

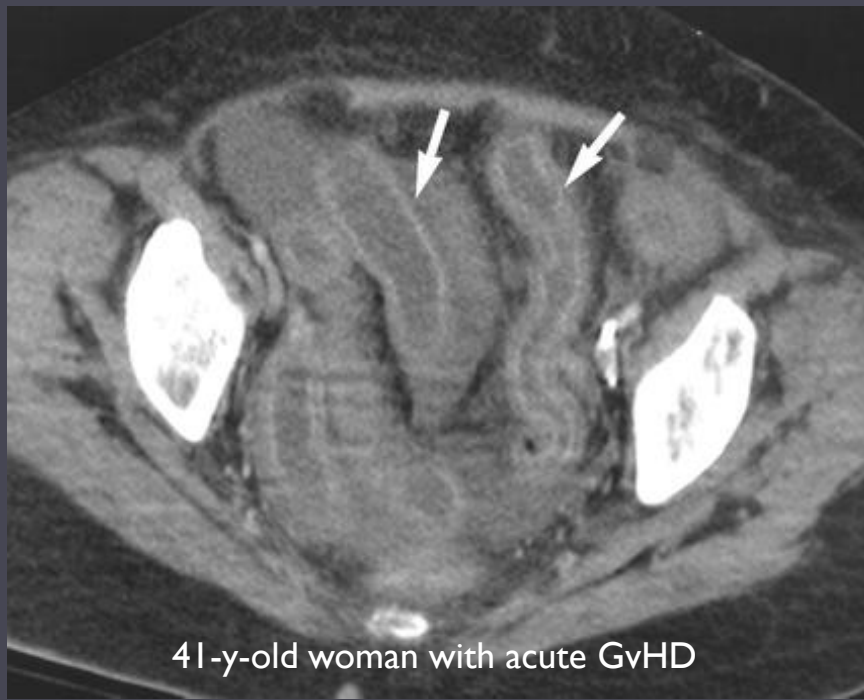
## CT Features with Pathologic Correlation of Acute Gastrointestinal Graft-Versus-Host Disease After Bone Marrow Transplantation in Adults

| <b>TABLE I</b> Intestinal CT Findings in Patients with Histologically Confirmed Acute Gastrointestinal Graft-Versus-Host Disease |     |     |
|----------------------------------------------------------------------------------------------------------------------------------|-----|-----|
| Finding                                                                                                                          | No. | %   |
| Wall thickening ( <i>n</i> = 22 patients)                                                                                        |     |     |
| Small-bowel                                                                                                                      | 22  | 100 |
| Large-bowel                                                                                                                      | 13  | 59  |
| Stomach                                                                                                                          | 2   | 9   |
| Distal esophagus                                                                                                                 | 2   | 9   |
| Enhancement <sup>a</sup> ( <i>n</i> = 13 patients)                                                                               |     |     |
| Mucosal                                                                                                                          | 7   | 54  |
| Serosal                                                                                                                          | 4   | 31  |
| Bowel-loop dilatation                                                                                                            | 5   | 23  |

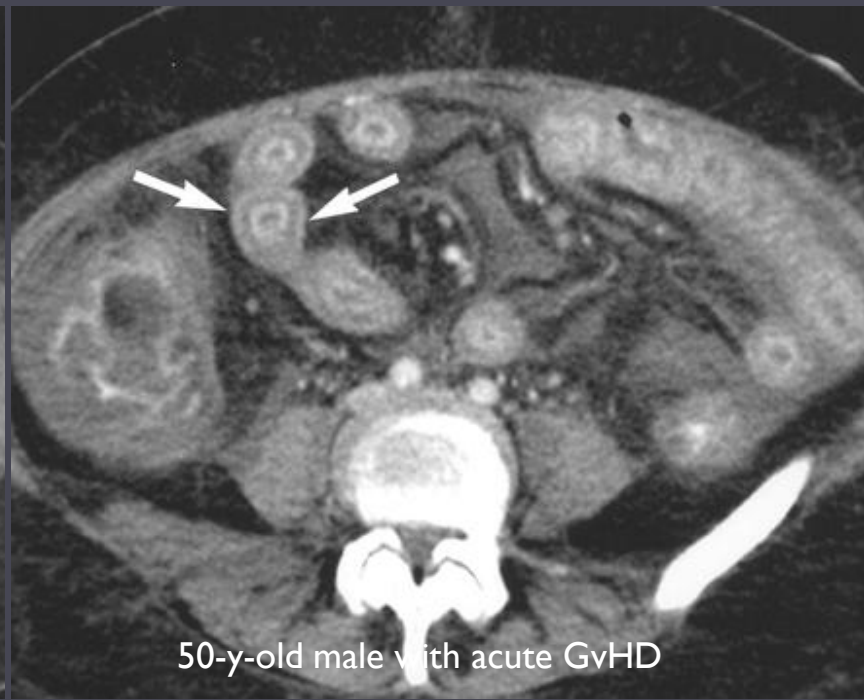
<sup>a</sup>Only 13 patients underwent contrast-enhanced CT.

Receveur de greffe de moelle osseuse. Douleurs abdominales et diarrhée.

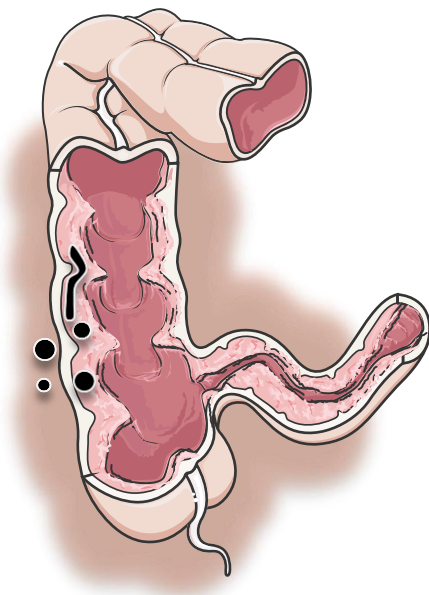




41-y-old woman with acute GvHD

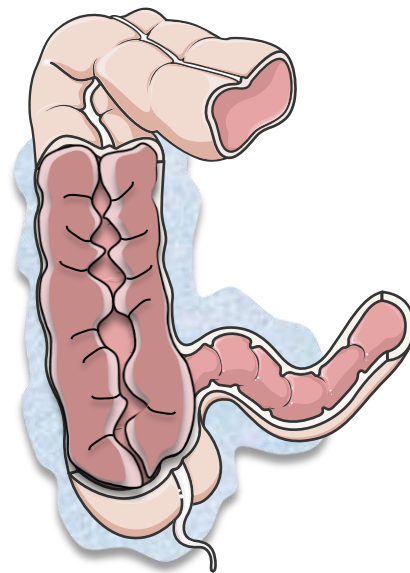


50-y-old male with acute GvHD



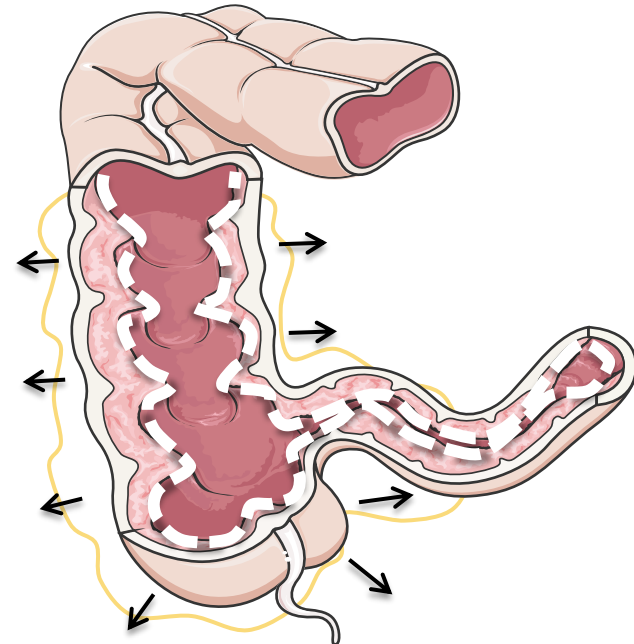
## Colite neutropénique

Epaississement pariétal  
Infiltration de la graisse  
**Pneumatose**



## Colite à *C. difficile*

Epaississement pariétal  
**'Signe de l'accordéon'**  
Ascite



## GvHD aiguë

Epaississement pariétal  
**Engorgement de Vasa recta**  
**Rehaussement muqueux**  
**Dilatation**

# Infections parenchymateuses

Principalement des micro-abcès, principalement foie et rate

# Groupes à risque pour les infections fongiques invasives chez les patients en cancérologie

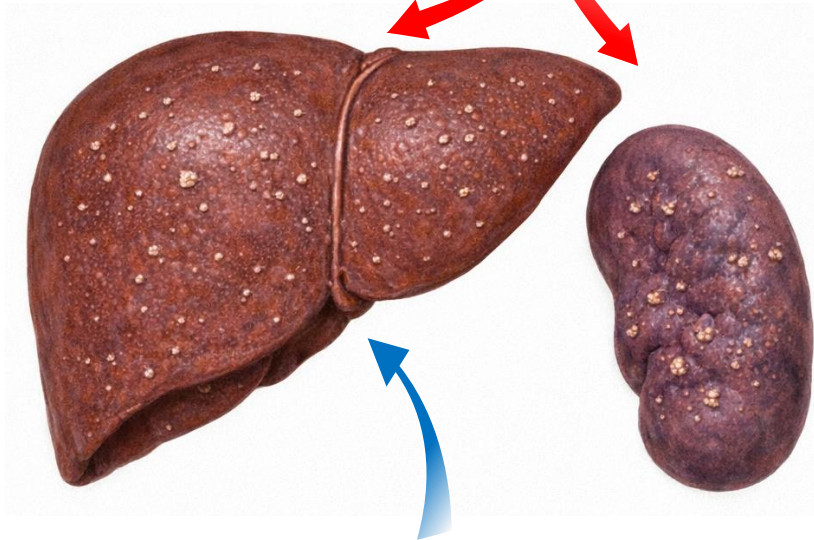


|                        |                                                                                                                                                                                                                                                                                  |
|------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Low risk               | Autologous bone marrow / stem cell transplant (SCT)<br>Childhood ALL<br>Lymphoma                                                                                                                                                                                                 |
| Intermediate low risk  | Moderate neutropenia 0.1-0.5 G/l < 3 weeks<br>Lymphocytes < 0.5 G/l + antibiotics<br>Older age • Central venous catheter                                                                                                                                                         |
| Intermediate high risk | Colonized > 1 site or heavy at one site<br><b>Neutropenia</b> < 0.5 to > 0.1 G/l, >3 to <5 weeks<br>AML • TBI • Allogenic matched sibling donor SCT                                                                                                                              |
| High risk              | <b>Neutropenia</b> <0.1 G/l, > 3 weeks OR <0.5 G/l > 5 weeks<br>Colonized by Candida tropicalis<br>Untreated or mismatched donor SCT • GVHD<br>Corticosteroids: >1mg/kg & neutroph. < 1 G/l > 1 week<br>Corticosteroids: >2mg/kg, > 1 week<br>High dose cytarabine • Fludarabine |

**Neutropenia (severity/duration), transplantation, immunosuppression**

# Infections fongiques → Micro-abcès

Porte d'entrée Vasculaire



Porte d'entrée Digestive

## Contamination

Sondes

Voies centrales

Plaies et cicatrices abdominale

Drains

Chirurgie

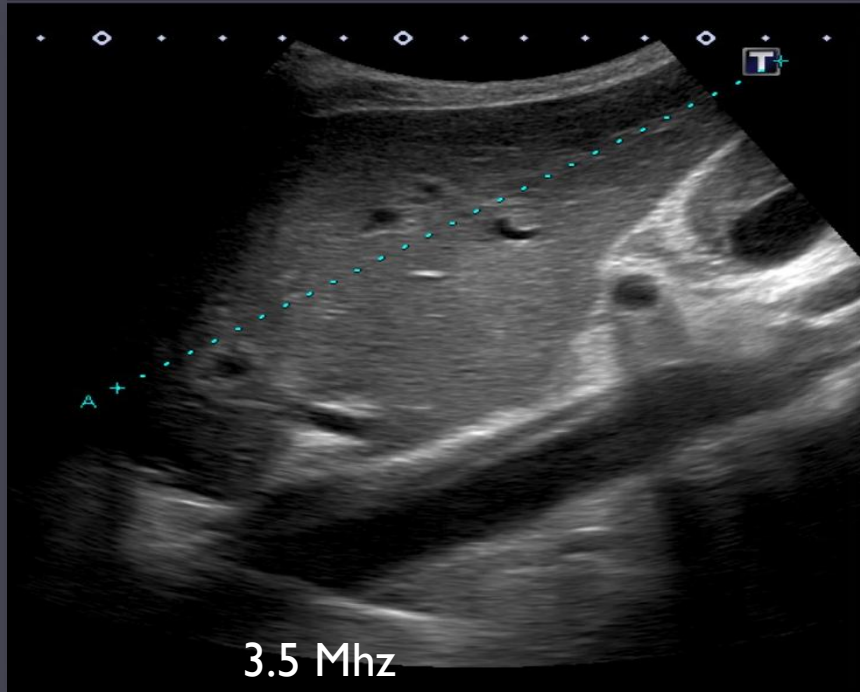
Nutrition parentérale

Ventilation mécanique

...

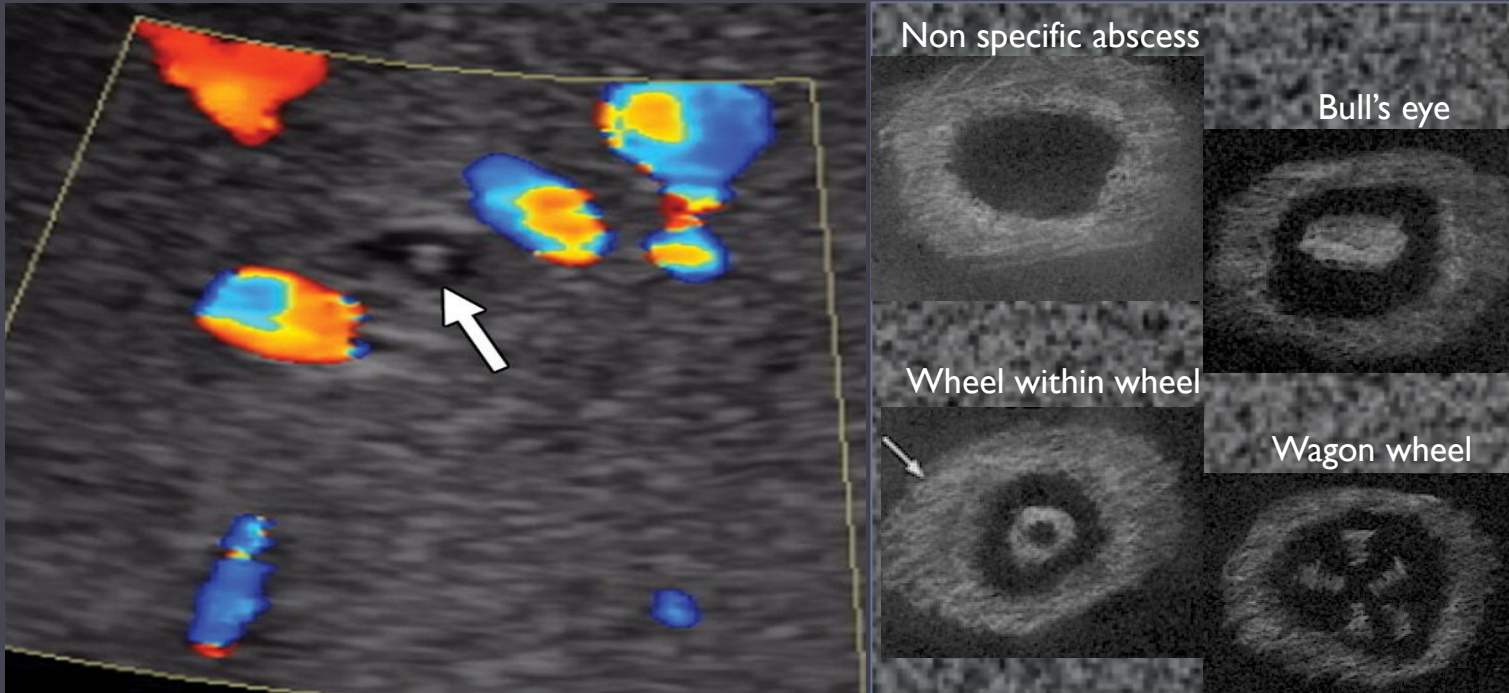
## Microabcès hépatiques

Immunodépression profonde post-induction d'une leucémie



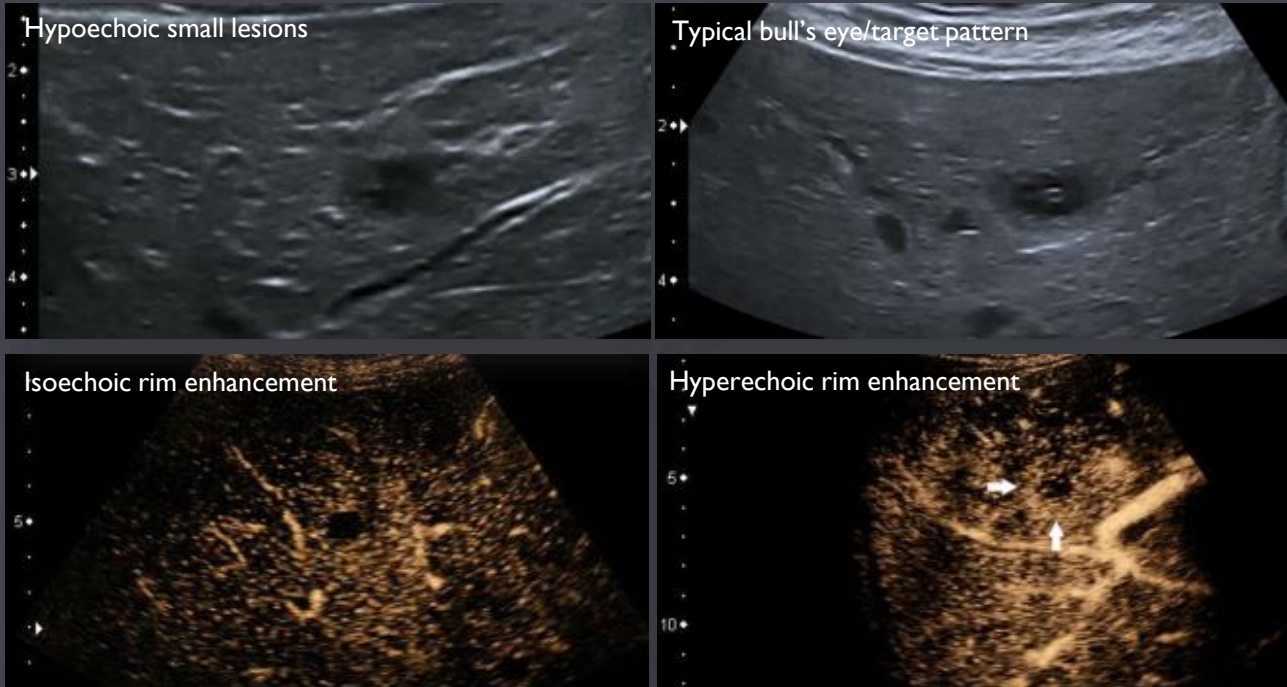
# Micro-abscès du foie et de la rate à l'échographie

Aspects les plus classiques

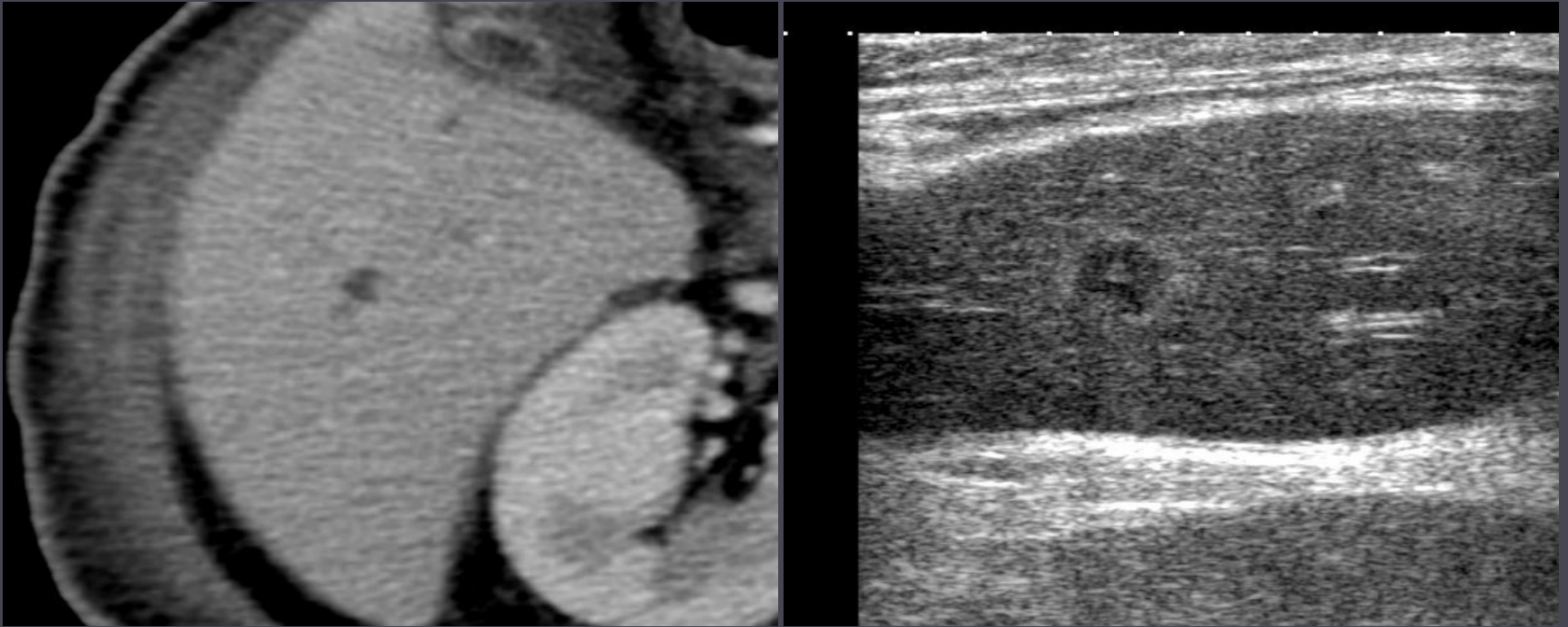


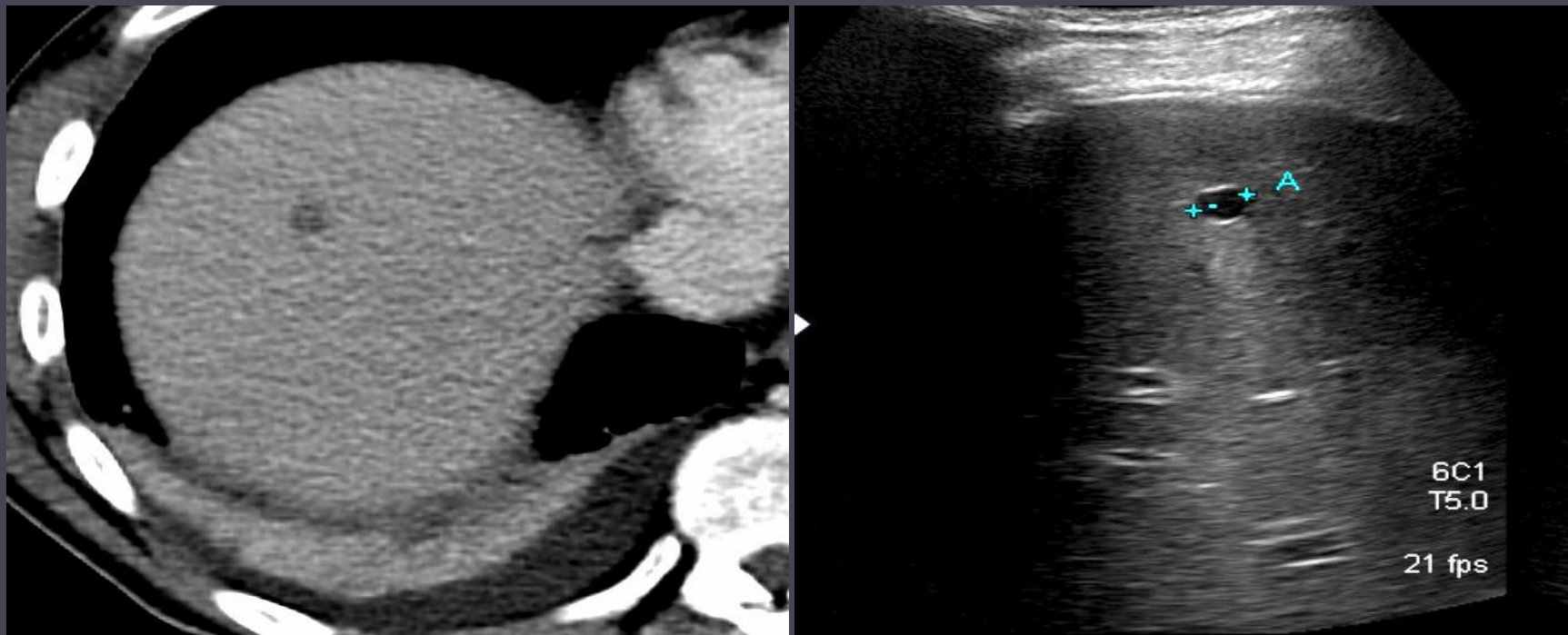
# Micro-abcès du foie et de la rate à l'échographie

## Candidose hépatique : aspects classiques



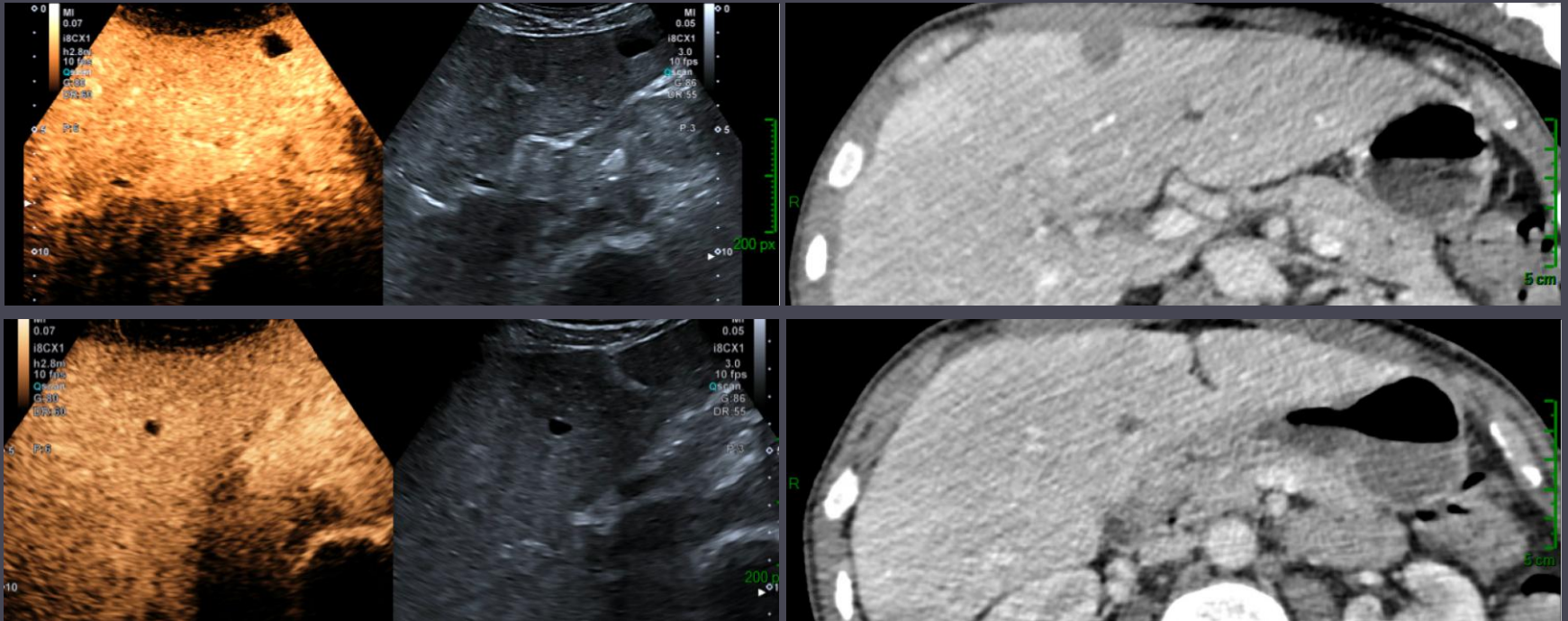
Bull's eye





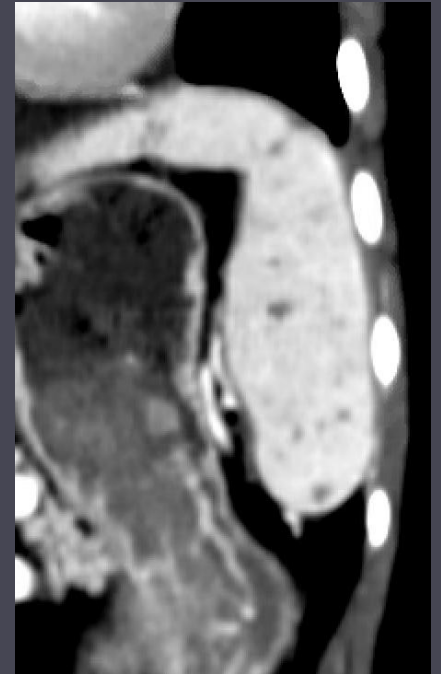
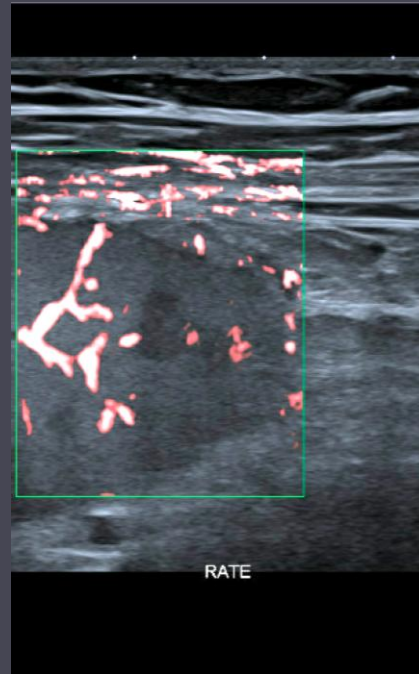
## Scanner versus US versus CE-US

VIH. Lymphome de Burkitt en cours de traitement. Candidose hépatique



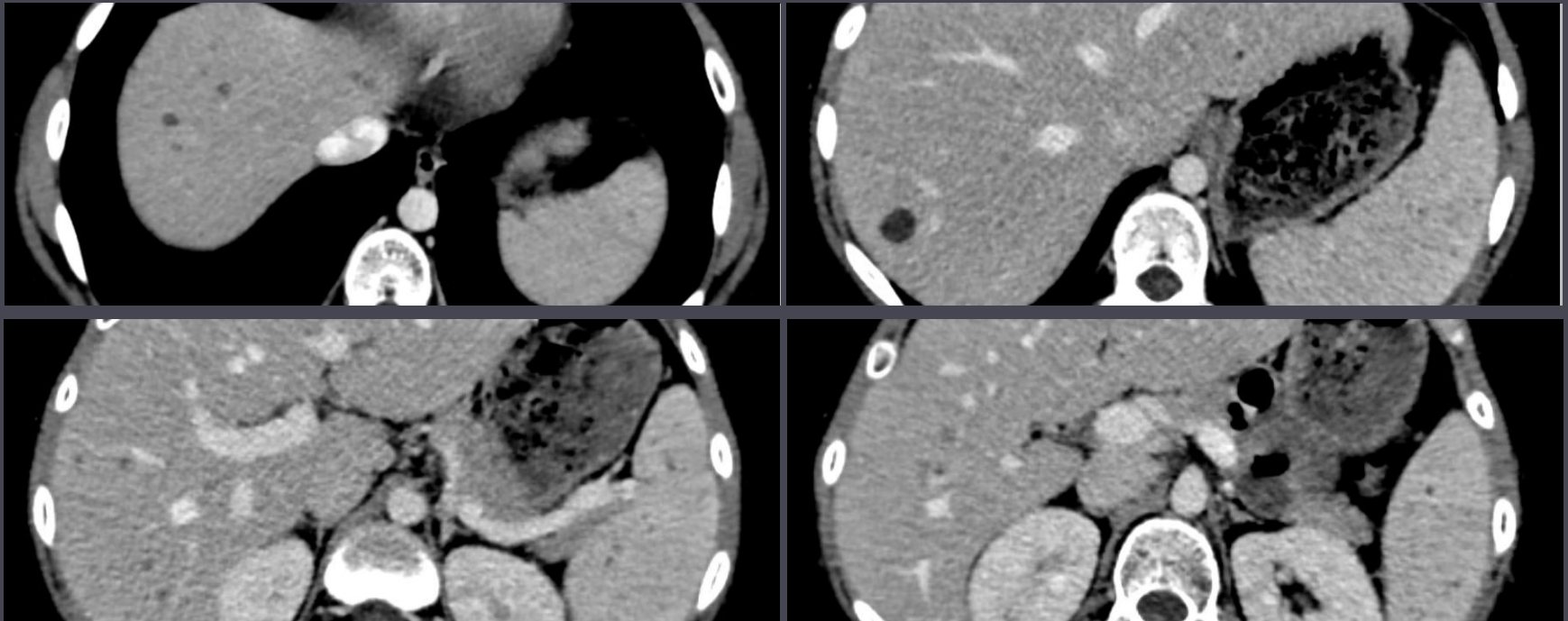
## Scanner versus US

20 ans. J13 d'une allogreffe pour LAM B. Candidose splénique



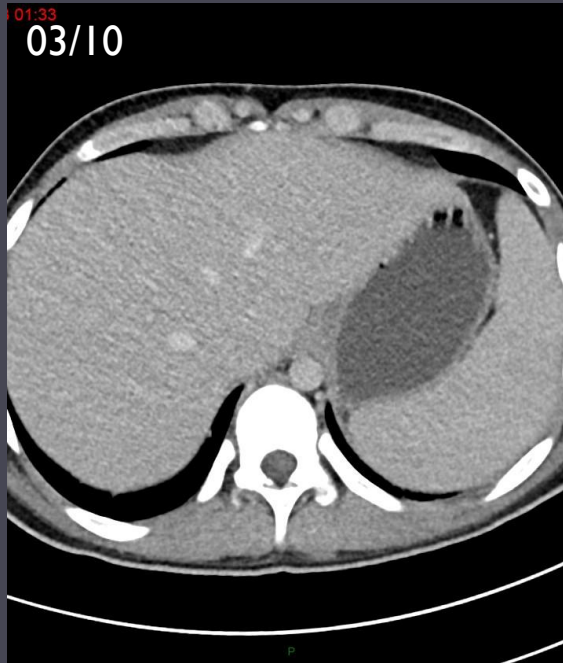
## Candidose hépato-splénique

18 ans, LAL T. Aplasie fébrile. Candidose hépato-splénique



LAL T en cours d'induction. Fièvre sans point d'appel

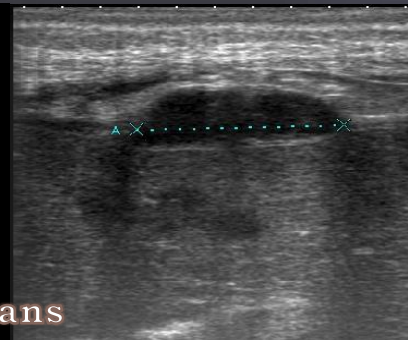
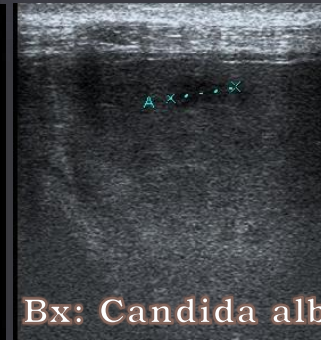
## Evolution hyper rapide !



Une cinétique inquiétante

61 ans, LAM, neutropénie profonde, fièvre sans point d'appel

## Evolution hyper rapide !

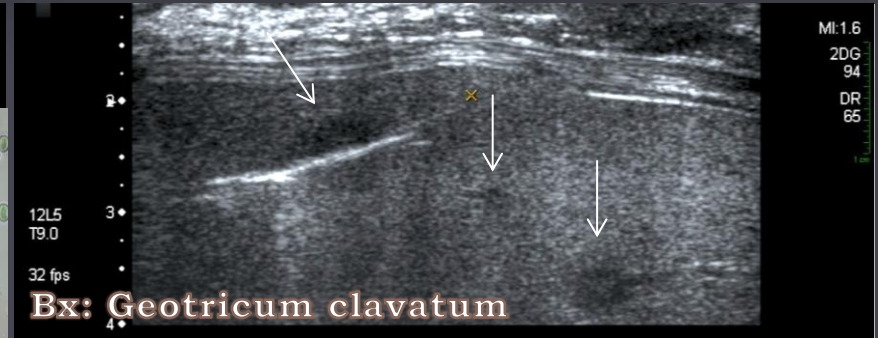
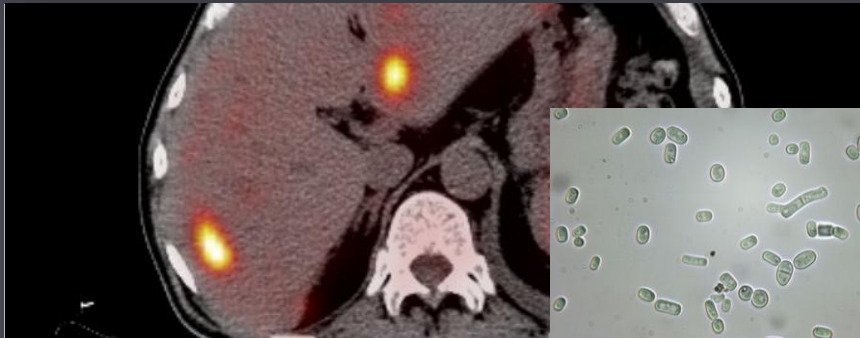


Bx: Candida albicans

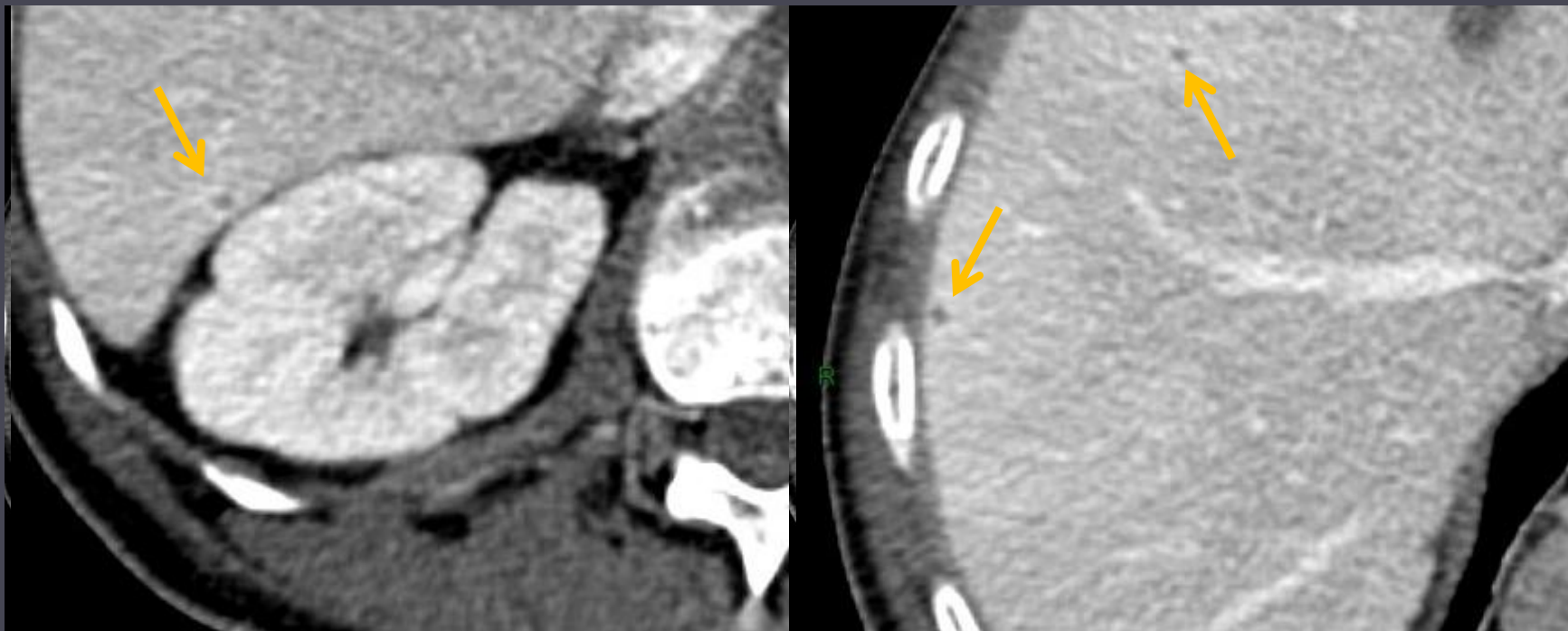


22 ans. LAM. Neutropénie fébrile.

## Evolution hyper rapide !

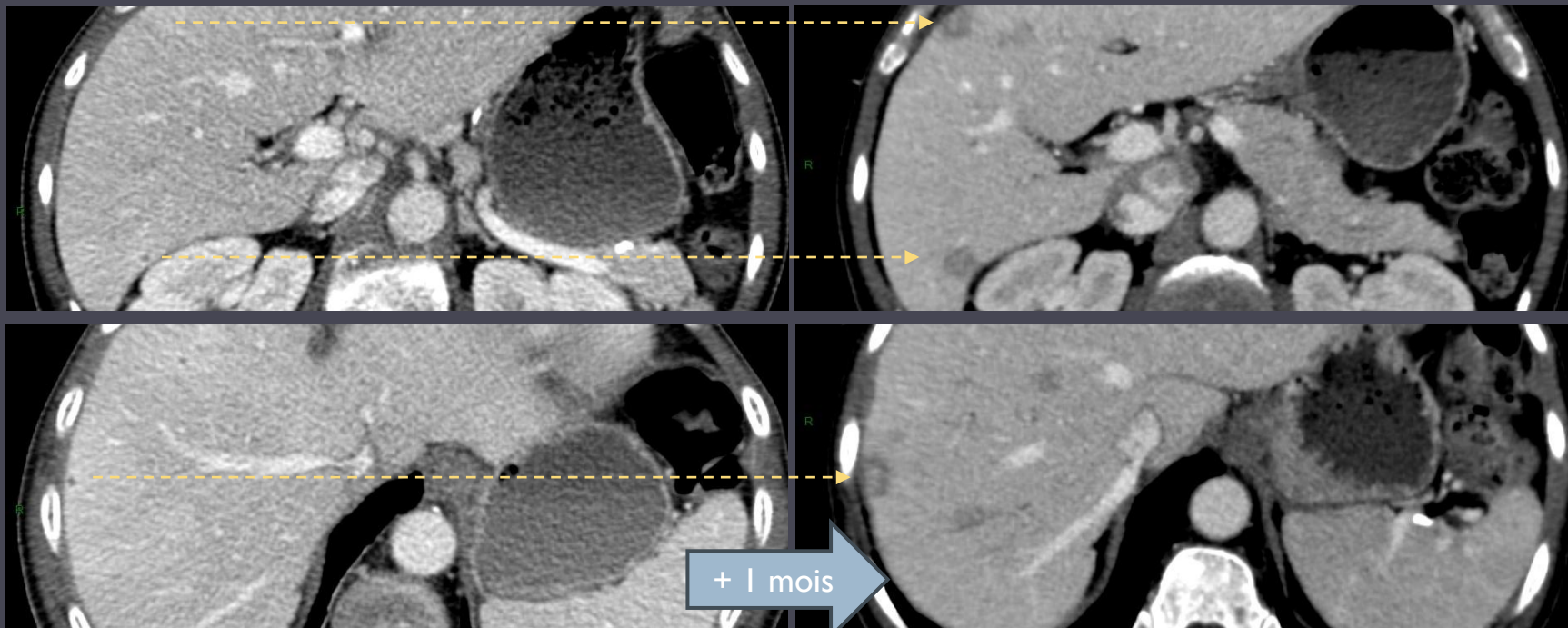


Candidose débutante chez une patiente traitée pour LAM. Fièvre à J30 après induction.

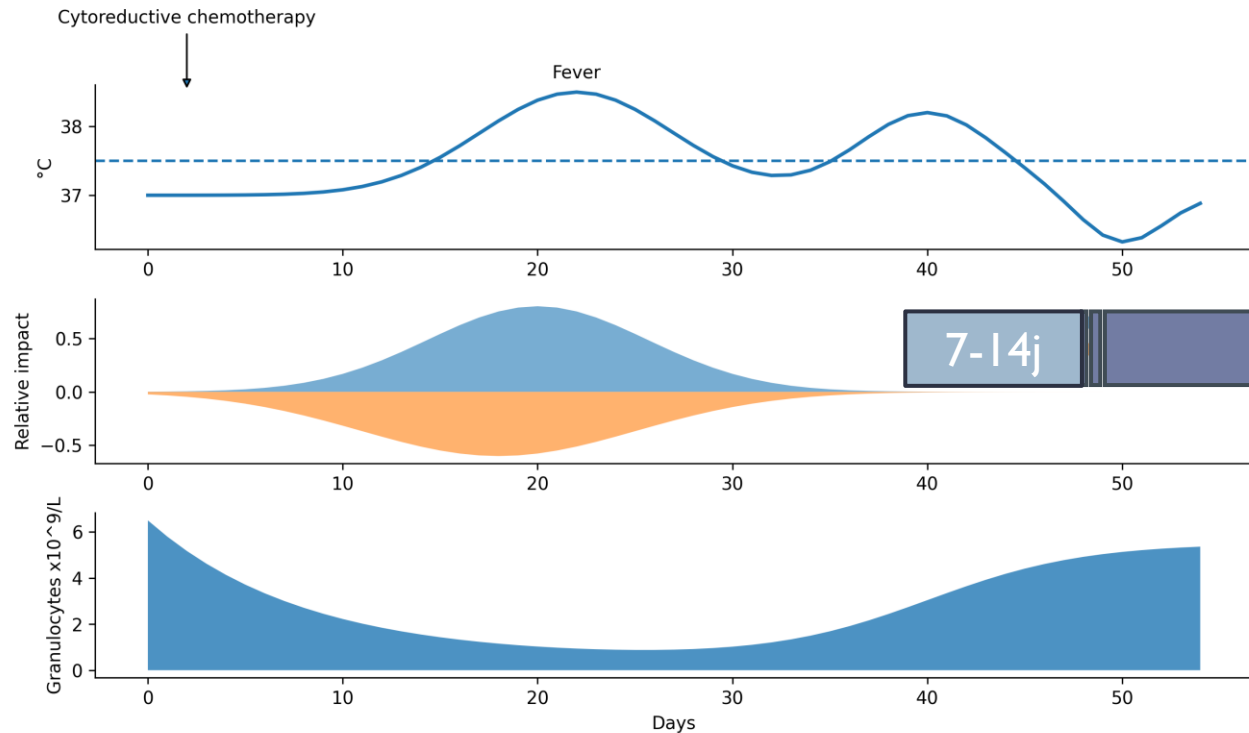


Contrôle 1 mois après traitement probabiliste par Ambisome

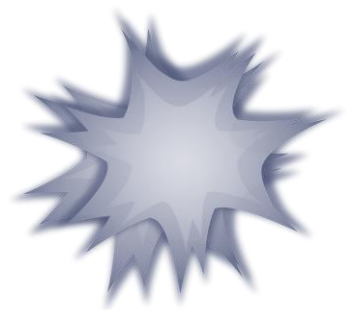
Le patient ne va pas mieux, le scanner non plus !



# Chimiothérapie → Neutropénie → IRIS



IRIS



# Syndrome Inflammatoire de Reconstitution Immunitaire (IRIS) post neutropénie fébrile

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- ▶ 7-14j après la repousse neutrophilique ( $>500/\text{mm}^3$ )
- ▶ Survient chez 10 à 30% des patients à haut risque
- ▶ Aggravation brutale chez des patients contrôlés sous prophylaxie/traitement
- ▶ **Mécanisme** : Libération massive de cytokines → Majoration inflammation locale → Simule une progression infectieuse
- ▶ Diagnostic : Clinique/Radio/Immuno + infection documentée. Bx si doute
- ▶ Traitement : Antifongiques adaptés + corticoïdes 1-2 mg/Kg/j

Ne pas arrêter l'antifongique !

RESEARCH ARTICLE

Open Access

# A systematic review of hepatic tuberculosis with considerations in human immunodeficiency virus co-infection

Andrew J Hickey<sup>1\*</sup>, Lilishia Gounder<sup>2,3</sup>, Mahomed-Yunus S Moosa<sup>2</sup> and Paul K Drain<sup>4,5</sup>

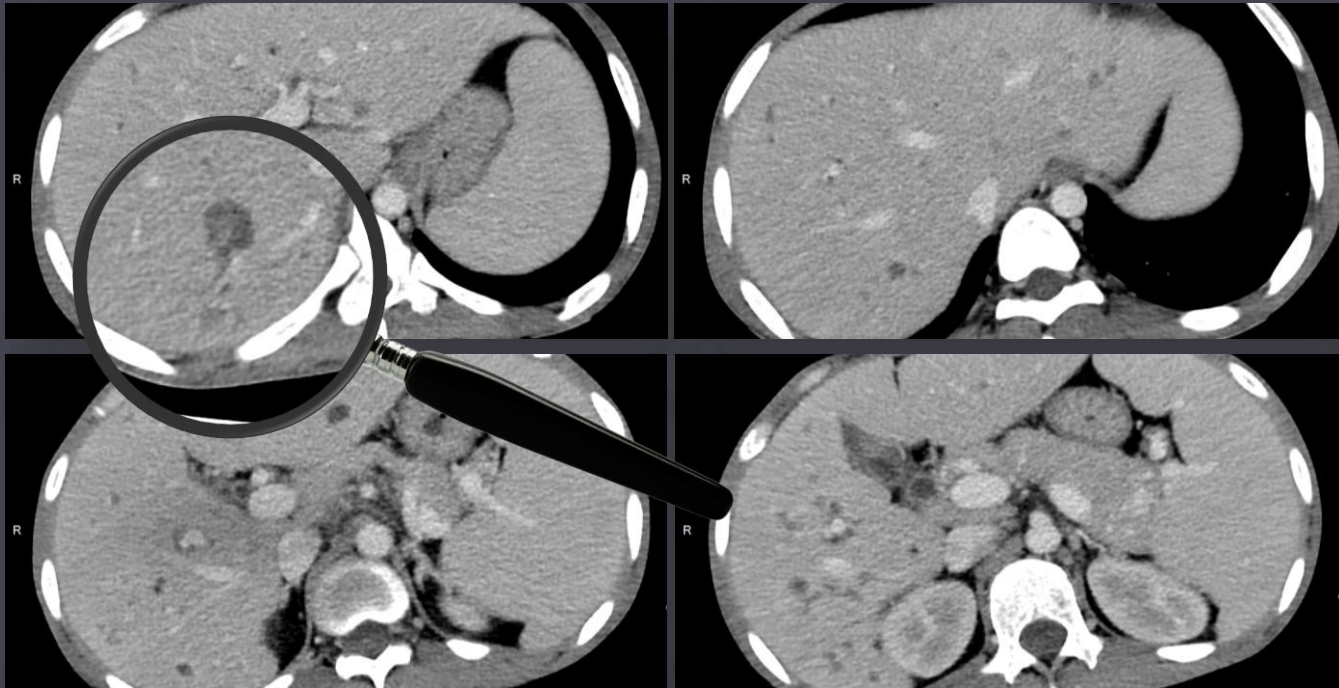
- ▶ Patients de regions endémiques
- ▶ Patients SIDA/HIV en rupture de traitement
- ▶ CT
  - ▶ Multiple micronoduleshypodenses (foie & rate)
  - ▶ Adénopathies fréquentes
- ▶ US
  - ▶ Micronodules hypoéchogènes
  - ▶ Parfois nodules échogènes

# Mycobactérioses and tuberculose

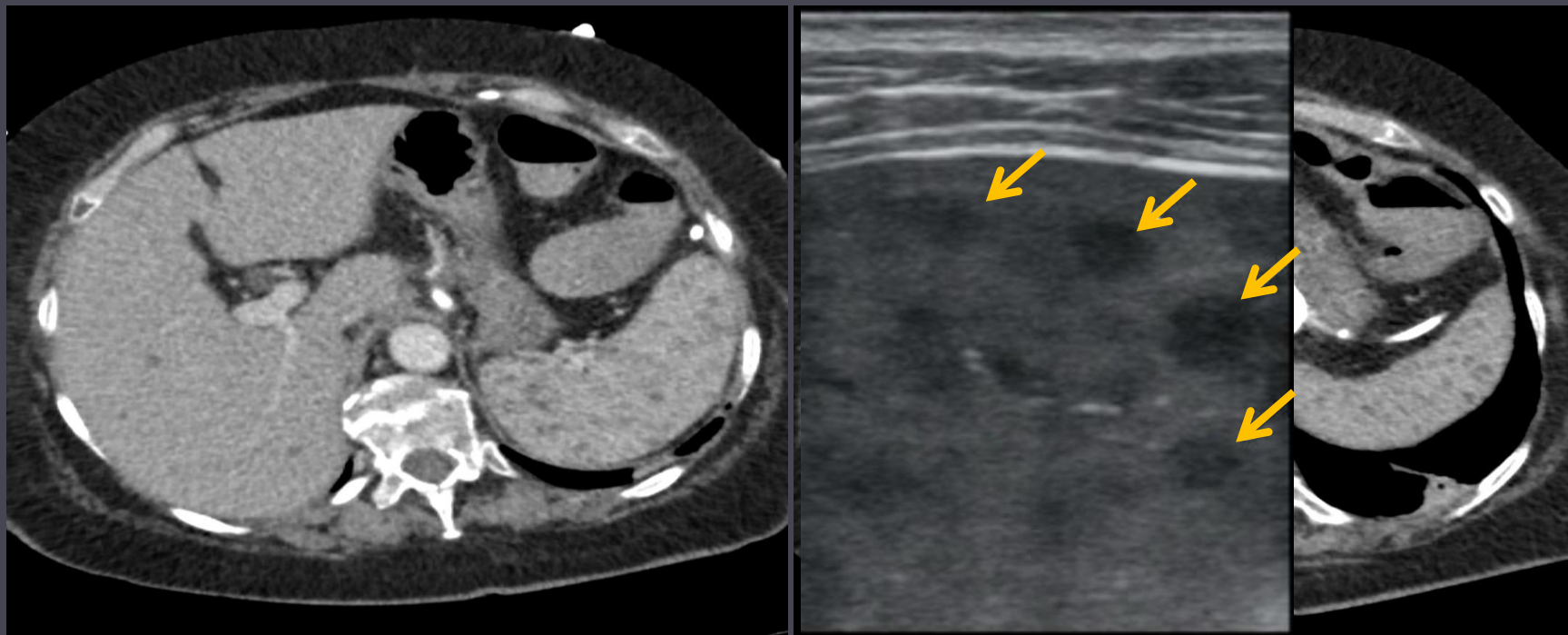


## Des granulomes plus que des microabcès

Patient HIV+. Immunodépression profonde. Nodules hépatiques. Tuberculose.



Patient. Au stade SIDA. Infection à *Mycobacterium avium*



# Conclusion

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- ▶ Importance du contexte clinique
  - ▶ Neutropénie
  - ▶ Délai après chimo/transplantation
  - ▶ SIDA
- ▶ Infections du tractus digestif : savoir reconnaître la colite neutropénique et la colite à clostridium principalement
- ▶ Infections des organes solides
  - ▶ Utiliser l'échographie (sonde haute fréquence) pour confirmation et biopsie
  - ▶ Attention à l'IRIS → Risque d'arrêter le traitement

