

Hepatic Abscess - Microbiology Full Clinical Guideline

Reference number: CG-MICRO/2019/002

Introduction

- One of the outcomes of:
 - o Microbial invasion of the liver parenchyma; and
 - The subsequent hepatic inflammatory response
 Is the formation of an encapsulated lesion containing necrotic immune cells and invading pathogens, i.e. a hepatic abscess.
- Hepatic abscesses can be caused by single pathogens (monomicrobial infection) or, more commonly, by multiple pathogens (polymicrobial infectious disease).
- Klebsiella pneumoniae and Escherichia coli are the most commonly diagnosed microbial causes of hepatic abscess.
- Bacteroides species, Enterococcus species, and the Streptococcus anginosus group (Streptococcus anginosus/constellatus/intermedius) are other relatively common bacterial causes.
- Less common causes include *Staphylococcus aureus* and *Streptococcus pyogenes*.
- The pathogens of hepatic abscesses are most commonly inoculated through a
 contiguous mechanism of transmission. Another focus of infection (e.g.
 cholangitis, cholecystitis) disseminates locally; the microorganism proliferates in
 the biliary tract and invades the liver parenchyma.
- Less commonly, inoculation is via a haematogenous mechanism of transmission. Infection of the portal vein 'pylephlebitis' (secondary to pancreatitis, appendicitis, diverticulitis, etc.) or bacteraemia (secondary to infective endocarditis, bowel cancer, etc.) can culminate in hepatic invasion.
- The pathogens of hepatic abscesses can also be inoculated via trauma, either penetrating or blunt.
- The purulent liver mass may manifest with localising symptoms and signs (e.g. abdominal pain, nausea, vomiting) and/or generalised stigmata of infectious disease (e.g. fever, chills, malaise).
- Temperatures > 38 ° C or < 36 ° C, a respiratory rate > 20 breaths/minute, a heart rate > 90 beats/minute, and hypotension can denote progression of localised infectious disease into sepsis and septic shock.

Differential diagnosis

- The symptoms and signs of hepatic abscess may overlap with the stigmata of other infective and non-infective pathologies:
 - o Infective:
 - Pneumonia, <u>acute cholecystitis</u>, <u>acute cholangitis</u>, viral hepatitis, and amoebic liver abscess.
 - NB Re amoebic liver abscess:
 - Risk factors include past medical/drug history of immunocompromise and social history of travel (e.g. refugee from/travellers to Africa [including South Africa], Asia [including India], North America [including Mexico], and South America).



- If the symptoms, signs, and/or pathology findings raise the differential diagnosis of Entamoeba histolytica hepatic abscess, please liaise with the hepatology, microbiology, and radiology teams regarding investigation and treatment.
- Non-infective:
 - Alcoholic hepatitis, drug-induced hepatitis, and liver cancer (primary or secondary).

Investigation

Radiology

- First line:
 - Ultrasound (US) of the liver.
- Second line:
 - Collaborate with the radiology consultant.

Microbiology

- With the range of bacterial pathogens, variations in bacterial resistance and susceptibility profiles, contraindications, side-effects, and with prolonged durations of weeks-months of antimicrobial chemotherapy, microbiological investigation enables best antibiotic practice:
 - Before starting antimicrobials:
 - Blood cultures x 2, drawn approximately 1-15 minutes apart, from 2 locations/venepunctures.
 - If radiology intervenes:
 - Pus for microscopy, culture, and susceptibilities (MC&S).
 Please notify the laboratory during the day or the microbiology biomedical scientist on call (via switchboard), if urgent MC&S of the radiological sample is required.

Blood sciences

Full blood count (FBC), C reactive protein (CRP), ± lactate, urea and electrolytes (U&Es), and liver function tests (LFTs).

Treatment

Radiological intervention

- Interventional radiology with:
 - o Percutaneous drainage via insertion of a pigtail catheter drain; or
 - o Percutaneous needle aspiration

Is the mainstay of hospital intervention for hepatic abscesses.

- Radiological intervention can be considered for uniloculated purulent liver masses of varying size, and also for multi-loculated hepatic abscesses.
- With regard to radiology:
 - Interventional radiology requires:
 - An electronic request; and
 - Informed consent for the procedure (https://derby.koha-ptfs.co.uk/cgi-bin/koha/opac-detail.pl?biblionumber=774); and
 - An up-to-date platelet count and clotting (https://derby.koha-ptfs.co.uk/cgi-bin/koha/opac-detail.pl?biblionumber=1577)

To be completed by the referring team.

 Please note, in general, local Trust policy requires omission of antiplatelets (e.g. clopidogrel for 5-7 days) and anticoagulants (e.g.



- warfarin for 5 days, apixaban or rivaroxaban for 48 hours) before radiological intervention.
- Possible exceptions wherein the clinical condition dictates drainage or antiplatelet/anticoagulant omission is contraindicated – require surgical consultant to interventional radiologist consultant discussion, regarding potential benefits and risks of intervention.

Surgical intervention

Rarely, surgical intervention can be contemplated. Uncommon scenarios raising
the possibility of drainage/resection in theatre – for example, hepatic abscess
rupture – require consultation with the senior hepatobiliary surgeon.

Empiric, intravenous antibiotics

- If the patient is clinically stable **and** for radiological intervention:
 - o Await drainage and withhold antimicrobial chemotherapy.
- If the patient has had drainage **or** if the patient is not for radiological intervention:
 - After blood cultures x 2:
 - If there are no clinical concerns regarding sepsis:

| First line | Piperacillin tazobactam 4.5 g 8 hourly |
|--|--|
| Second line, if non-immediate without | Ceftriaxone 2 g 24 hourly and |
| systemic involvement penicillin allergy | Metronidazole 500 mg 8 hourly |
| Third line, if immediate rapidly evolving or | Ciprofloxacin 400 mg 12 hourly and |
| non-immediate with systemic involvement | Metronidazole 500 mg 8 hourly |
| penicillin allergy | |

If there are clinical concerns regarding sepsis (life threatening organ dysfunction caused by a dysregulated host immune response to infection) secondary to hepatic abscess:

| First line | Piperacillin tazobactam 4.5 g 6 hourly ± If there are clinical concerns regarding the risk of methicillin resistant <i>Staphylococcus aureus</i> (MRSA), vancomycin or teicoplanin, <u>dose as per hospital guidelines</u> , vancomycin target pre dose level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l |
|---|--|
| Second line, if non-immediate without systemic involvement penicillin allergy | Ceftazidime 2 g 8 hourly and Vancomycin or teicoplanin, <u>dose as per</u> <u>hospital guidelines</u> , vancomycin target pre dose level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Metronidazole 500 mg 8 hourly |
| Third line, if immediate rapidly evolving or non-immediate with systemic involvement penicillin allergy | Ciprofloxacin 400 mg 8 hourly and Vancomycin or teicoplanin, dose as per hospital quidelines, vancomycin target pre dose level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Metronidazole 500 mg 8 hourly |



Directed antibiotics (with susceptibilities)

• Intravenous, according to susceptibilities:

| | ing to susceptibilities: | | |
|---------------------------------------|--------------------------------|---|--|
| | First line | Second line, <u>if</u> | Third line, <u>if immediate</u> |
| | | non-immediate | rapidly evolving or non- |
| | | without systemic | immediate with systemic |
| | | involvement | involvement penicillin |
| | | penicillin allergy | <u>allergy</u> |
| Escherichia coli | Narrowest | Ceftriaxone 2 g | Ciprofloxacin 400 mg 12 |
| | spectrum of | 24 hourly | hourly (consider per oral |
| | amoxicillin or co- | • | [absorption 60-80% and |
| | amoxiclav or | | peak concentration in |
| | piperacillin | | bile/peak concentration |
| | tazobactam | | in serum 2800-4500%]) |
| | standard dosage | | |
| Klebsiella | Narrowest | Ceftriaxone 2 g | Ciprofloxacin 400 mg 12 |
| pneumoniae | spectrum of co- | 24 hourly | hourly (consider per oral |
| pricarriorilae | amoxiclav or | 24 Hourry | [absorption 60-80% and |
| | piperacillin | | peak concentration in |
| | tazobactam | | bile/peak concentration |
| | | | • |
| Danianaidan Futanan | standard dosage | | in serum 2800-4500%]) |
| <u>-</u> | | • | cially – can be associated |
| with polymicrobial hepa | | | |
| | | | coccus species and also |
| uncultured Gram negat | | | |
| Bacteroides/anaerobe | Co-amoxiclav 1.2 | Ceftriaxone 2 g | Ciprofloxacin 400 mg 12 |
| | g 8 hourly | 24 hourly and | hourly and |
| | | Metronidazole | Metronidazole 500 mg 8 |
| | | 500 mg 8 hourly | hourly |
| Enterococcus species | Co-amoxiclav 1.2 | Vancomycin or | Vancomycin or |
| | g 8 hourly | teicoplanin, dose | teicoplanin, dose as per |
| | | as per hospital | hospital guidelines, |
| | | guidelines, | vancomycin target pre |
| | | vancomycin | dose level 15-20 mg/l, |
| | | target pre dose | |
| | | i laidel bie dose | i telcoblanin tardet bre - i |
| | | | teicoplanin target pre |
| | | level 15-20 mg/l, | dose level 15-30 mg/l |
| | | level 15-20 mg/l, teicoplanin target | dose level 15-30 mg/l and |
| | | level 15-20 mg/l, teicoplanin target pre dose level | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 |
| | | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and |
| | | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ceftriaxone 2 g | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and Metronidazole 500 mg 8 |
| | | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ceftriaxone 2 g 24 hourly and | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and |
| | | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ceftriaxone 2 g 24 hourly and Metronidazole | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and Metronidazole 500 mg 8 |
| Strontos a saus | Co. americles A.C. | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ceftriaxone 2 g 24 hourly and Metronidazole 500 mg 8 hourly | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and Metronidazole 500 mg 8 hourly |
| Streptococcus | Co-amoxiclav 1.2 | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ceftriaxone 2 g 24 hourly and Metronidazole 500 mg 8 hourly Ceftriaxone 2 g | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and Metronidazole 500 mg 8 hourly Vancomycin or |
| Streptococcus anginosus group | Co-amoxiclav 1.2 g 8 hourly | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ceftriaxone 2 g 24 hourly and Metronidazole 500 mg 8 hourly Ceftriaxone 2 g 24 hourly and | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and Metronidazole 500 mg 8 hourly Vancomycin or teicoplanin, dose as per |
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| - | | level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ceftriaxone 2 g 24 hourly and Metronidazole 500 mg 8 hourly Ceftriaxone 2 g 24 hourly and Metronidazole | dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and Metronidazole 500 mg 8 hourly Vancomycin or teicoplanin, dose as per hospital guidelines, vancomycin target pre dose level 15-20 mg/l, teicoplanin target pre dose level 15-30 mg/l and Ciprofloxacin 400 mg 12 hourly and |



400 mg 8 hourly

- After 7-14 days of intravenous antimicrobial chemotherapy, if the patient is afebrile, observations stable, and inflammatory markers downward trending, collaborate with the physician/surgeon regarding their preference for: (1) per oral step down; or (2) outpatient parenteral antimicrobial therapy (OPAT).
- After 7-14 days of intravenous antimicrobial chemotherapy, if the patient is febrile, observations unstable, and/or inflammatory markers upward trending, collaborate with the physician/surgeon and radiologists regarding investigation/intervention, update the microbiologist, and continue intravenous therapy.

• Per oral, according to susceptibilities:

| Per oral, according to st | • | <u> </u> | |
|---|----------------------|------------------------|-------------------------|
| | First line | Second line | Third line |
| Escherichia coli | Narrowest spectrum | Ciprofloxacin | Co-trimoxazole |
| | of amoxicillin 1 g 8 | 500 mg 12 | 960 mg 12 hourly |
| | hourly; or co- | hourly | |
| | amoxiclav 625 mg 8 | | |
| | hourly plus | | |
| | amoxicillin 500 mg 8 | | |
| | hourly | | |
| Klebsiella pneumoniae | Co-amoxiclav 625 | Ciprofloxacin | Co-trimoxazole |
| | mg 8 hourly plus | 500 mg 12 | 960 mg 12 hourly |
| | amoxicillin 500 mg 8 | hourly | |
| | hourly | | |
| Bacteroides, Enterococcus | | | |
| with polymicrobial hepatic | | | |
| encompassing the culture | | | species and also |
| uncultured Gram negatives | | example: | <u> </u> |
| Bacteroides/anaerobe | Co-amoxiclav 625 | <u>Ciprofloxacin</u> | Co-trimoxazole |
| | mg 8 hourly plus | 500 mg 12 | 960 mg 12 hourly |
| | amoxicillin 500 mg 8 | hourly and | and |
| | hourly | Metronidazole | Metronidazole |
| | | 400 mg 8 hourly | 400 mg 8 hourly |
| Enterococcus species | Co-amoxiclav 625 | Linezolid 600 mg | Linezolid 600 mg |
| | mg 8 hourly plus | per oral 12 | per oral 12 |
| | amoxicillin 500 mg 8 | hourly and | hourly and |
| | hourly | <u>Ciprofloxacin</u> | Co-trimoxazole |
| | | 500 mg 12 | 960 mg 12 hourly |
| | | hourly and | and |
| | | Metronidazole | Metronidazole |
| | | 400 mg 8 hourly | 400 mg 8 hourly |
| Streptococcus anginosus | Co-amoxiclav 625 | Clindamycin 300 | Linezolid 600 mg |
| group | mg 8 hourly plus | mg 6 hourly and | 12 hourly and |
| | amoxicillin 500 mg 8 | <u>Ciprofloxacin</u> | Ciprofloxacin 500 |
| | hourly | 500 mg 12 | mg 12 hourly |
| | | hourly | and |
| | | | Metronidazole |



Directed, outpatient parenteral antimicrobial therapy

Collaborate with the OPAT consultant.

Empiric, per oral or outpatient parenteral antimicrobial therapy

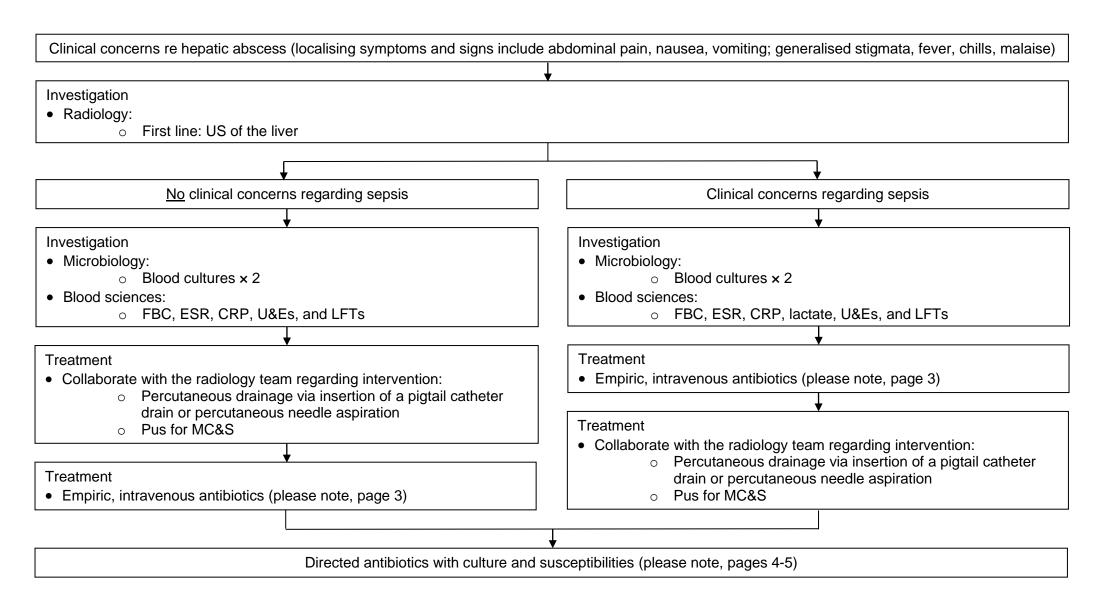
 If there are symptoms, signs, and/or radiological features of hepatic abscess, and the microbiology is negative, collaborate with a microbiologist regarding empiric options.

Duration of antibiotics

- Before discharge to the community, medical/surgical team to collaborate with radiology regarding ± re-imaging.
- If for per oral step down or OPAT, monitor bloods (FBC, CRP, U&Es, and LFTs) weekly-fortnightly.
- Courses of antibiotics 2-6 weeks.
- If radiology have intervened and if the patient is afebrile, observations stable, and inflammatory markers have resolved:
 - o 2-4 weeks.
- If surgery have intervened and if the patient is afebrile, observations stable, and inflammatory markers have resolved:
 - o 2 weeks.
- If neither radiology nor surgery intervene:
 - o 6 weeks.
- Follow up with the medical/surgical team on intravenous or per oral therapy.



Management





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Document control

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