

KISS: Clostridioides Difficile

NICE NG199 July 2021 - CKS 2023

Background:

- C. Diff. overgrowth does not cause infection per se, it is the **toxin production that causes the diarrhoea**; this is important when considering testing (a 2-step process to identify both the organism and the toxin) see below:
 - C. Diff. carriage is not uncommon so C. Diff. organism +ve but -ve toxin does not necessarily mean infection is present.
- Awareness of the drivers of C. diff. (in particular broad-spectrum antibiotics) has improved dramatically over the past 15 years, resulting in decreases in C. Diff.
 - Cases/100,000 population dropped from 107/100,000 in 2007/08 to 25/100,000 in 2013/14, however, rate of decline has slowed since, and 2021/22 saw a slight increase in cases.

Assessment:

- Risk factors? Age >65; broad-spectrum antibiotics recently; concurrent antibiotics or long courses; previous C. Diff. (recurrence rate ~20% after the first episode); exposure to known case; underlying morbidity; drugs (e.g. PPIs).
- Assess severity: Consider bloods to help assess severity (FBC, U&E)
 - Non-severe WCC <15, Cr rise \leq 50% of baseline, temp \leq 38.5oC
 - Severe = one of WCC \geq 15, Cr rise > 50% of baseline, temp >38.5oC
 - Fulminant/life threatening shock, hypotension, toxic megacolon or perforation, rapid clinical deterioration.

Management:

- Decisions on admission will be dictated by clinic judgement taking into account the severity of symptoms and underlying risk factors (e.g. age, support at home, ?bloody diarrhoea, comorbidities, risks of dehydration and ability to take oral antibiotics) +/- WCC/renal function, but if any severe features present admit.
- Testing:
 - Send stool sample immediately if C. Diff. suspected/at risk.
 - Do not re-test if +ve test if still symptomatic within the same episode.
 - Only re-test to confirm recurrent C. Diff. if symptoms resolve then recur.
 - Do not test to confirm cure people can remain C. Diff. +ve even after successful treatment.



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Management Continued:

- **Review existing antibiotic use** stop unless essential if essential can it be switched to lower risk abx?
- **Review medications** and stop if possible PPIs, laxatives, and SADMAN meds (e.g. NSAIDs, diuretics, ACEi, SGLT2i).
- **Treatment:** (Consider prompt specialist advice prior to treatment, or if empiric tx considered whilst awaiting test result)
 - **1st line:** Vancomycin 125mg orally QDS for 10 days.
 - **2nd line:** Fidaxomicin 200mg orally BD for 10 days and if ongoing symptoms seek specialist advice.
 - **Relapse*** (further C. Diff infection within 12 weeks) = fidoxamicin (dose above).
 - **Recurrence*** (further C. Diff. infection after 12 weeks) = vancomycin or fidoxamicin (doses as above).
 - But if relapse or recurrence* low threshold for seeking microbiology advice.
- Advise on maintaining hydration, preventing spread, and safety net.
- Warn patients on **natural history** diarrhoea should resolve in 1-2 weeks, but usually unable to determine if treatment effective before day 7. **Reassess** if symptoms do not respond as expected or worsen.
- Consider referral for **faecal microbiota transplant** for recurrent C. Diff. infection if ≥2 previous episodes.
- Prebiotics or probiotics?
 - NICE advise people taking abx not to take prebiotics or probiotics to prevent C. diff. infection.
 - NICE found <u>1 meta-analysis</u> showing probiotics statistically significantly reduce the incidence of C. difficile infection in inpatients but not in outpatients or those in mixed settings.
 - There is some evidence for probiotics in reducing abx associated diarrhoea in general, just not specifically for C. Diff:
 - <u>Cochrane</u> found high-dose probiotics ARE effective at reducing antibiotic-associated diarrhoea in children, NNT = 9, using *Lactobacillus rhamnosus* or *Saccharomyces boulardii* - there is uncertainty about other types of probiotic.
 - <u>Meta-analysis published in J Clin Gastroenterology 2021</u> showed a relative risk reduction in antibiotic-associated diarrhoea of 38% when probiotics were taken early in antibiotic treatment.