







Plasma Cell Infiltration on Histopathological Samples of Chronic Bone and Joint Infections due to Cutibacterium acnes : A series of 21 cases

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- Histopathological definition of bone and joint infections (BJIs):
 - Mirra's criterion : ≥ 5 neutrophils/field in 5 High power Fields (HPF; x400)
- This definition is validate for acute BJIs only
- This threshold seems too high for the diagnosis of chronic BJIs
 - Especially due to *Cutibacterium acnes* and coagulase negative staphylococci
- Plasma cell is the prototype of the cell recruited in chronic infections

Parvizi J et al. J Arthroplasty. 2018. Bori G et al. Biomed Res Int. 2018. Bori G et al. Modern Pathol. 2006. Tande AJ et al. Clin Microbiol Rev. 2014. Kashima TG et al. Virchows Arch. 2015.



• Primary objective:

 To confirm that Mirra's criterion is not adequate for the diagnosis of chronic C. acnes BJIs

• <u>Secondary objective</u>:

• To determine if **plasma cell inflammation** could be a useful criterion for histopathological diagnosis of **chronic BJIs du to** *C. acnes*



- 25 consecutive patients from 2009 to 2013 with *C. acnes BJI* were selected from Lyon BJI cohort study (CRIOAc Lyon)
- Histological analysis was performed only on 21 patients with at least 2 *C. acnes* positive monobacterial cultures on separate deep samples

Introduction Objectives Methods Results Discussion & Conclusion

Histopathological analysis

- **High neutrophil infiltration** (Mirra's criterion : ≥ 5/field in 5 HPFs)
- High plasma-cell infiltration (CRIOAc Lyon's criterion : ≥ 5/field in 5 HPFs)
- Low neutrophil infiltration (<2, 2-5/field in 5 HPFs)
- Low plasma-cell infiltration (<2, 2-5/field in 5 HPFs)
- Other criteria : necrosis, granulation tissue, giant cells, fibrin

Patients were defined as infected if high infiltration (neutrophil or plasma-cell) was present

Introduction

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Clinical data of patients with *C. acnes* BJI

| | Total population, n=25 |
|--|------------------------|
| Gender, n (%) | |
| Female | 11 (44%) |
| Male | 14 (56%) |
| Median age, years (IQR) | 61.5 (40.5-75) |
| Median delay between previous surgery and current diagnosis, days (IQR) | 322 (179-786) |
| Type of implant | |
| Prosthetic joint, n (%) | 16 (64%) |
| Osteosynthesis, n (%) | 9 (36%) |
| Type of surgery | |
| Debridement and implant removal, n (%) | 22 (88%) |
| Debridement and implant retention, n (%) | 3 (12%) |

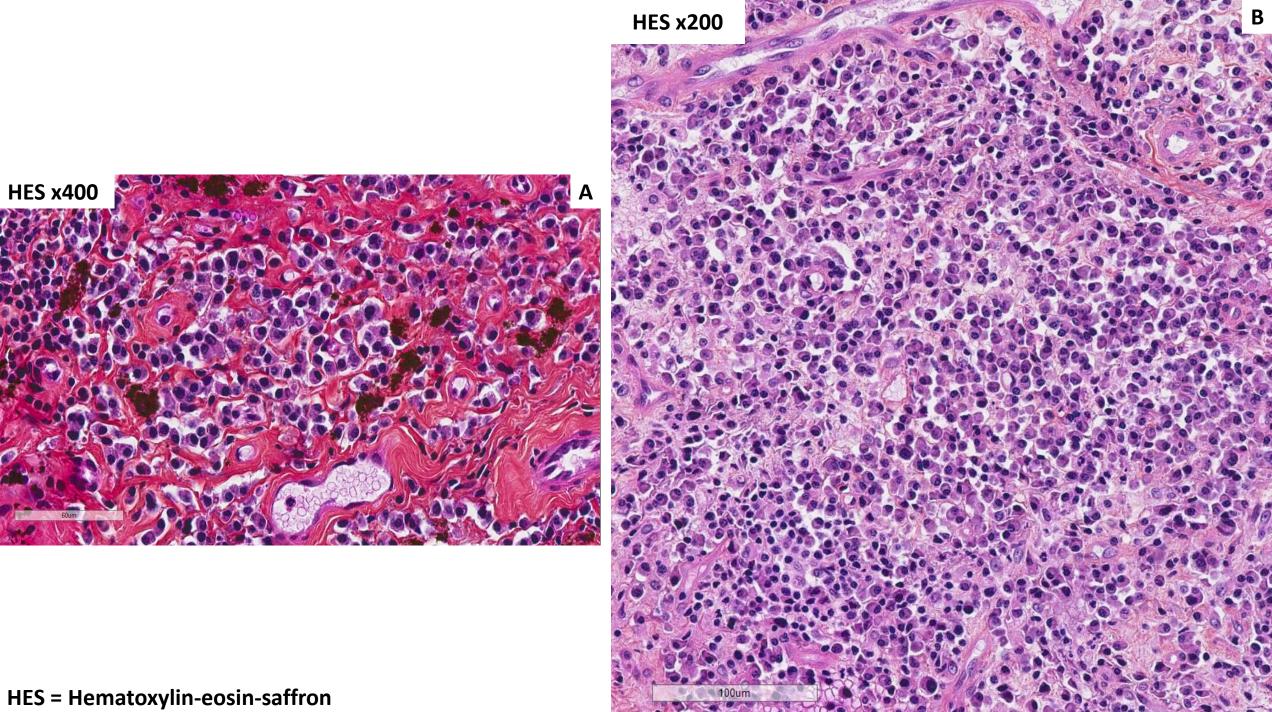
| | Methods | Results |
|--|---|---------|
| Criteria | % of patient | |
| Mirra's criterion (\geq 5 neutrophils/field in 5 HPFs) | 57.1% (12/21) | |
| CRIOAc Lyon's criterion (≥ 5 Plasma cells/field in 5 HPFs) | 71.4% (15/21) | |
| Mirra's and/or CRIOAC Lyon's criteria | <mark>81%</mark> (17/21) | |
| Neutrophil inflammation 2-5/field in 5 HPFs <2/field in 5 HPFs No PMN infiltration | 4.8% (1/21) 14.3% (3/21) 23.9% (5/21) | |
| Plasma cell inflammation 2-5/field in 5 HPFs <2/field in 5 HPFs No plasma-cell inflammation | 23.9% (5/21) 4.8% (1/21) 0% (0/21) | |
| Fibrine | 76.2% (16/21) | |
| Necrosis | 9.5% (2/21) | |
| Granulation tissue | 62% (13/21) | |
| Giant cells | 33.3% (7/21) | |

| | Methods | Results Discussion & Conclusion |
|--|---|--|
| Criteria | % of patient | |
| Mirra's criterion (\geq 5 neutrophils/field in 5 HPFs) | 57.1% (12/21) | →Adding CRIOAc Lyon's criterion had |
| CRIOAc Lyon's criterion (≥ 5 Plasma cells/field in 5 HPFs) | 71.4% (15/21) | restored 5/21 histopathological diagnosis of BJIs (23.9%) |
| Mirra's and/or CRIOAC Lyon's criteria | <mark>81%</mark> (17/21) | |
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| Plasma cell inflammation 2-5/field in 5 HPFs <2/field in 5 HPFs No plasma-cell inflammation | 23.9% (5/21) 4.8% (1/21) 0% (0/21) | |
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|--|---|--|--|
| Criteria | % of patient | | |
| Mirra's criterion (\geq 5 neutrophils/field in 5 HPFs) | 57.1% (12/21) | \rightarrow In 9/21 cases with negative Mirra's | |
| CRIOAc Lyon's criterion (≥ 5 Plasma cells/field in 5 HPFs) | 71.4% (15/21) | criterion (42.9%): - CRIOAc Lyon criterion +: 55.6% (5/9) | |
| Mirra's and/or CRIOAC Lyon's criteria | 81% (17/21) | - 2-5 plasma cells/field +: 44.4% (4/9) | |
| Neutrophil inflammation 2-5/field in 5 HPFs <2/field in 5 HPFs No PMN infiltration | 4.8% (1/21) 14.3% (3/21) 23.9% (5/21) | Fibrine +: 66.7% (6/9) Granulation tissue +: 44.4% (4/9) Necrosis +: 11.1% (1/9) Giant cells +: 33.3% (3/9) | |
| Plasma cell inflammation 2-5/field in 5 HPFs <2/field in 5 HPFs No plasma-cell inflammation | 23.9% (5/21) 4.8% (1/21) 0% (0/21) | | |
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| | | | Results | Discussion & Conclusion |
|--|--|--|---------|----------------------------|
|--|--|--|---------|----------------------------|

| | Osteosynthesis implant (n=9) | Prosthetic implant (n=12) |
|---|------------------------------|---------------------------|
| Mirra's criterion (≥ 5 PMNs/field in 5 HPFs) | 44.4 % (4/9) | 66.7% (8/12) |
| CRIOAc Lyon's criterion (≥ 5 Plasma cells/field in 5 HPFs) | <mark>88.9%</mark> (8/9) | 58.3% (7/12) |



HES = Hematoxylin-eosin-saffron

- This is the first study to describe plasma cell inflammation in chronic BJIs
- In our study:
 - **CRIOAc Lyon's criterion** appeared to be a **better diagnostic criterion** than Mirra's criterion (71.4% *versus* 57.1%)
 - adding CRIOAc Lyon's criterion have restored histopathological diagnosis in 23.9% (5/21 cases)
- However, it is a small series and plasma-cells are not specific of chronic BJIs

 → Plasma cells could evoke an infection due to *C. acnes*, but must be part of a multidisciplinary approach (biological and clinical criteria).

Bori G et al. Biomed Res Int. 2018. Kashima TG et al. Virchows Arch. 2015. Bori G et al. Modern Pathol. 2006. Pace T et al. J Arthroplasty. 1997. Plasma Cell Infiltration on Histopathological Samples of Chronic Bone and Joint Infections due to Cutibacterium acnes : A series of 21 cases

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Thank you for your attention

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