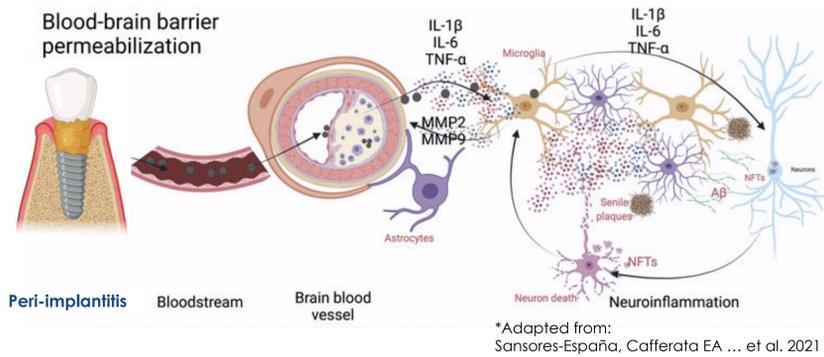


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Background

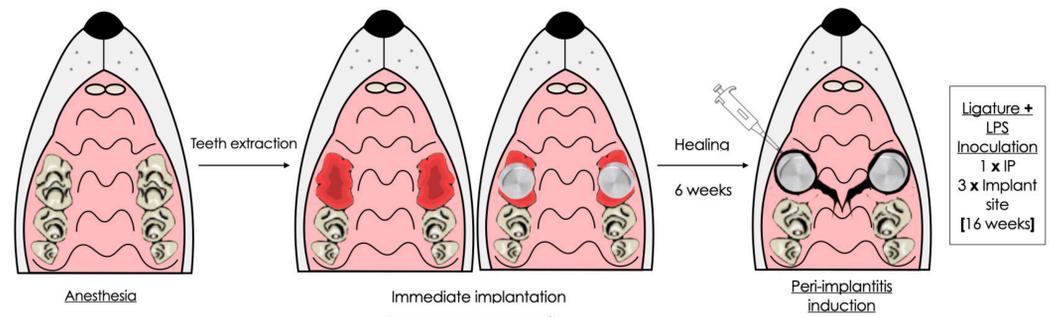


Hypothesis: Chronic inflammation and dysbiosis during peri-implantitis can cause systemic inflammation and bacteremia, which are closely associated to neuroinflammation that leads to the onset of neurodegenerative pathologies like Alzheimer's disease. However, there is **no evidence yet**.

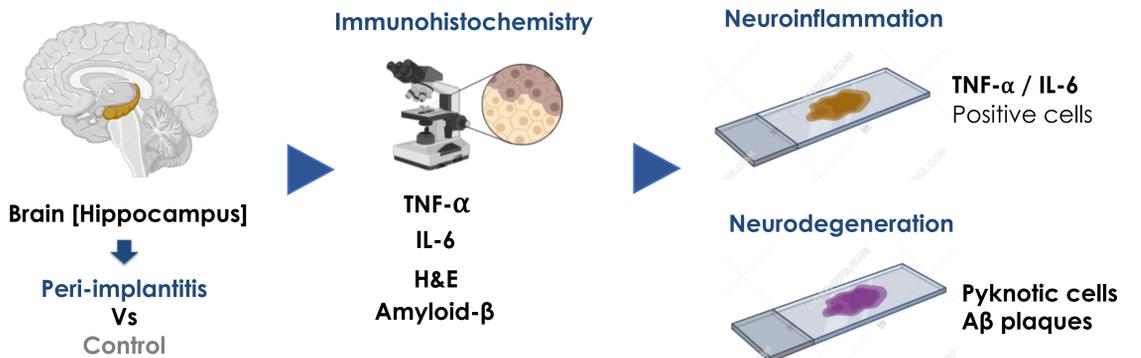
Aim: This study aimed to investigate the possible **association** between ligature-induced PI and **neuropathological changes in the brain** in a rat model

Methods

1. Chronic peri-implantitis lesion model



2. Cerebral tissue analyses



Results

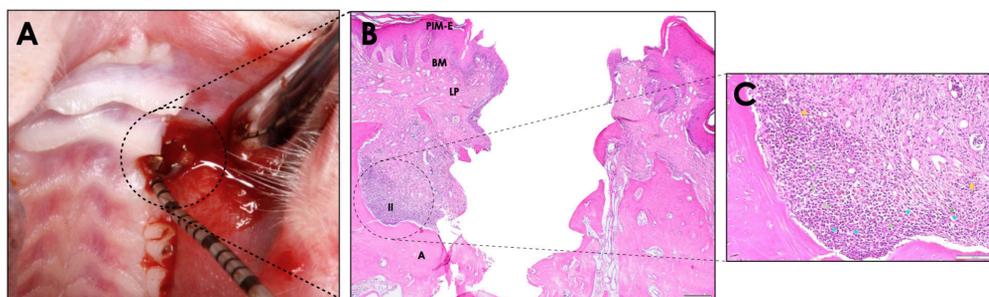


Figure 1. Experimental peri-implantitis. **A)** Clinical signs of inflammation were evidenced by the presence of bleeding on probing and suppuration at ligature removal. **B)** Representative H&E staining of an experimental peri-implantitis lesion [10x]. **C)** [20x] augmentation on **B** showing PI lesion inflammatory infiltrate, with the presence of polymorphonuclears, monocytes and some lymphocytes. PIM-E: peri-implant mucosa epithelium; BM: basal membrane; LP: lamina propria; II: inflammatory infiltrate; AB: alveolar bone.

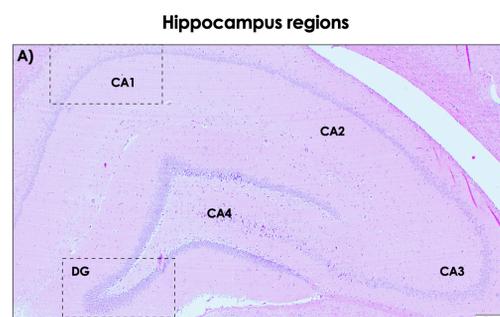


Figure 2A. Hippocampus showing cellular morphological alterations following experimental PI. **A)** Representative image showing the hippocampus regions -CA1, CA2, CA3, CA4 and DG-, the regions of interest CA1 and DG are delimited by dashed squares [5x]

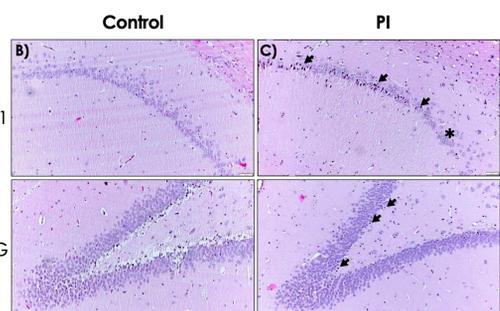


Figure 2B and C. Hippocampus showing cellular morphological alterations following experimental PI. **B)** Representative images of the CA1 and DG hippocampus regions of the control group [10x] and **C)** the peri-implantitis group, showing morphologically altered neurons and glial cells [10x]. CA: Cornu ammonis; DG: dentate gyrus; PI: peri-implantitis group; i: pyramidal neuron; →: astrocyte.

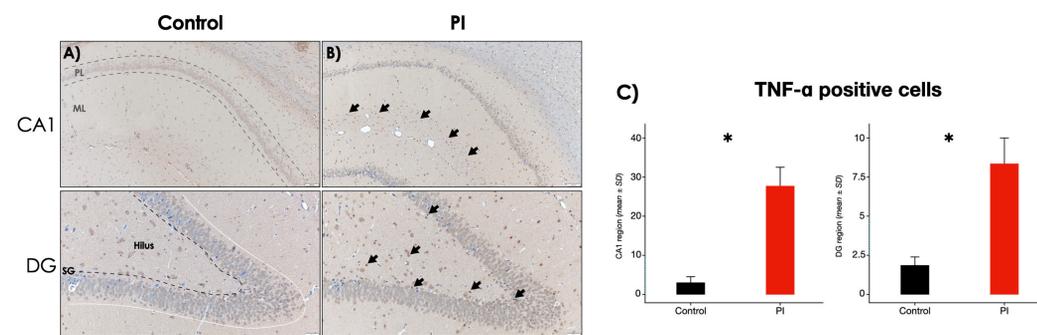


Figure 3. Hippocampus TNF-α immunostaining. **(A, B)** Representative hippocampus images showing the CA1 and DG regions [10x]. **(C)** Relative quantification of TNF-α + cells in the CA1 and DG regions of the hippocampus. *p < 0.05. PL: pyramidal layer; ML: molecular layer; SG: subgranular cell zone; G: granular cell zone; →: TNF-α positive cell.

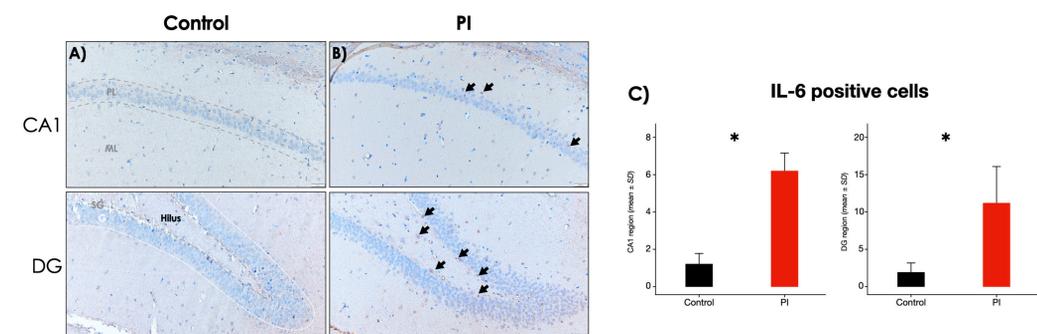


Figure 4. Hippocampus IL-6 immunostaining. **(A, B)** Representative hippocampus images showing the CA1 and DG regions [10x]. **(C)** Relative quantification of IL-6+ cells in the CA1 and DG regions of the hippocampus. *p < 0.05. PL: pyramidal layer; ML: molecular layer; SG: subgranular cell zone; G: granular cell zone; →: IL-6 positive cell.

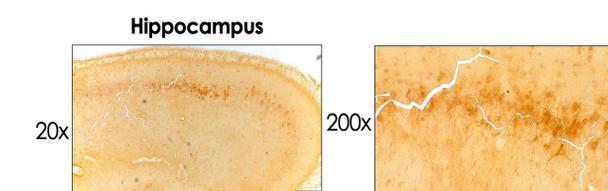


Figure 5. Representative hippocampus Aβ immunostaining. No Aβ positive immunostaining could be detected in any of the specimens.

Conclusion

➤ Within the limitations of the present experimental exploratory study, our analysis showed that chronic peri-implantitis lesions led to the increased detection of IL-6 and TNF-α in the hippocampus, showing neuroinflammatory signs associated with neurodegenerative disorders, such as Alzheimer's disease.