

SALIVA AND URINE METABOLOME IN KIDNEY TRANSPLANT RECIPIENTS.

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INTRODUCTION & AIM

Metabolome represents the collection of low molecular weight compounds that originate from metabolic pathways of the body, drugs, and microorganisms in the oral cavity. Small molecules extracted from tissues or detected in body fluids are measured using mass spectrometry. Targeted approaches using quantitative assays and validated by LC-MS/MS have indicated that intermediates of the transmethylation pathway, as well as arachidonic acid-derived bioactive metabolites, might be predictive markers for kidney transplant rejection. Metabolomics analysis of oral cavity samples facilitates differentiation between patients with periodontitis and healthy individuals. **The aim of this study was to evaluate the biofluid metabolome of kidney transplant recipients and their possible association with periodontitis.**

METHOD

- ✓ Cross-sectional, analytical, and descriptive study;
- ✓ Four groups;
- ✓ Urine and saliva collections;
- ✓ Gas Chromatography-Mass Spectrometry (GC-MS/MS);
- ✓ Statistical programs: Statistica 12, SPSS 20, Past 3.0, Origin Pro 2019, and MetaboAnalys 5.0

Inclusion criteria: (i) kidney transplantation; (ii) diagnosis of periodontitis; (iii) sites presenting probing pocket depth (PPD) ≥ 4 mm; (iv) those who underwent immunosuppressive therapy; and (v) GC without systemic changes and without the use of medications.

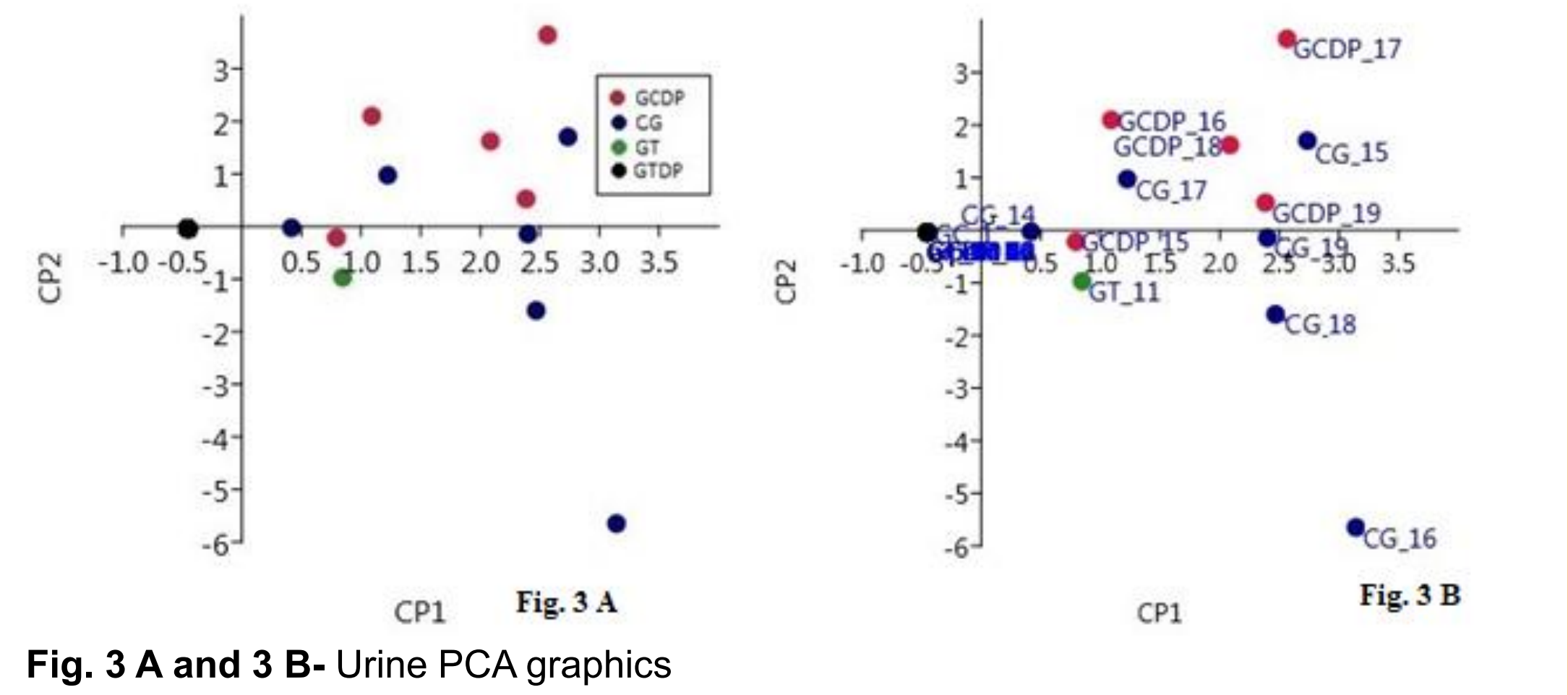
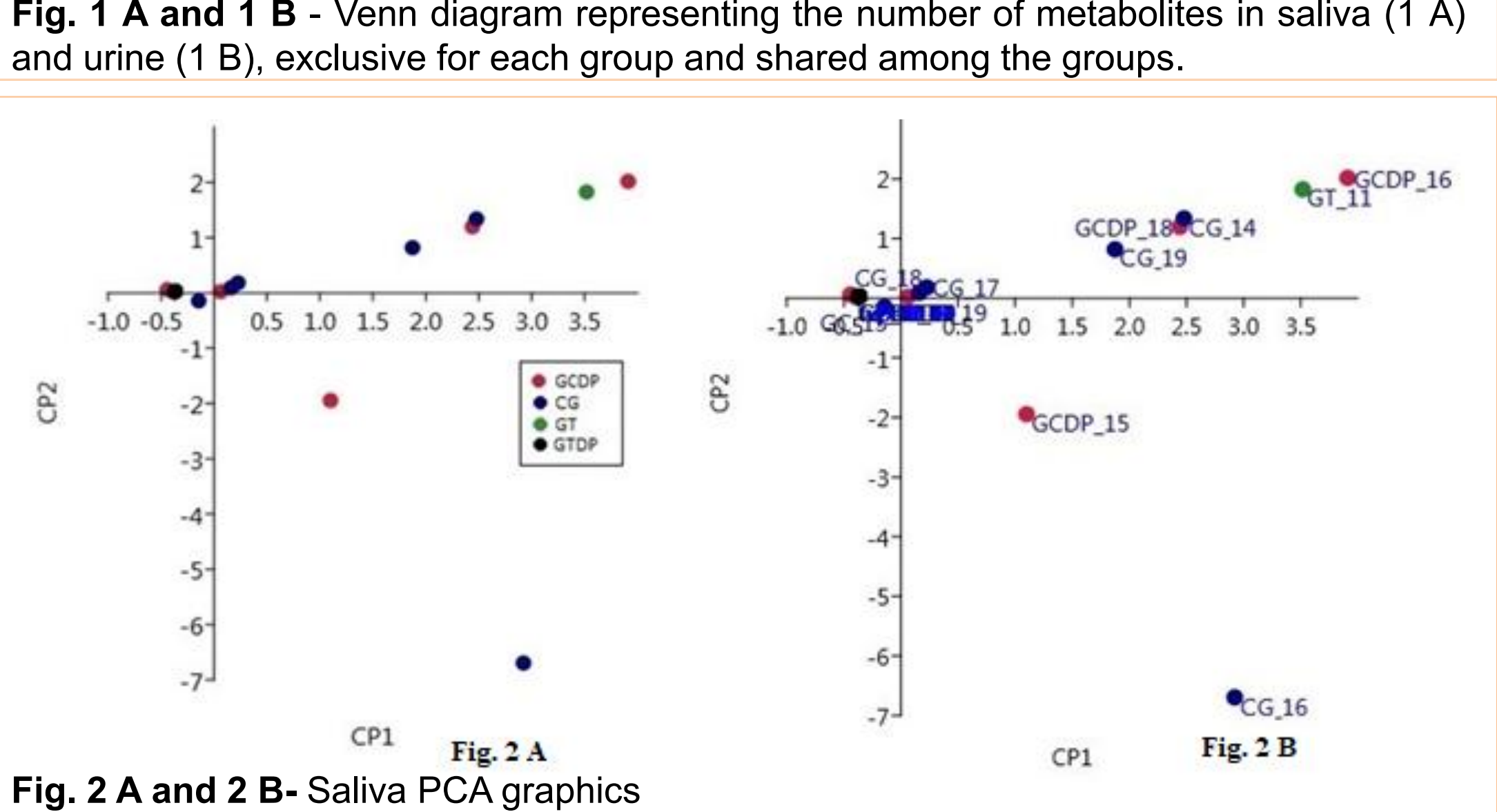
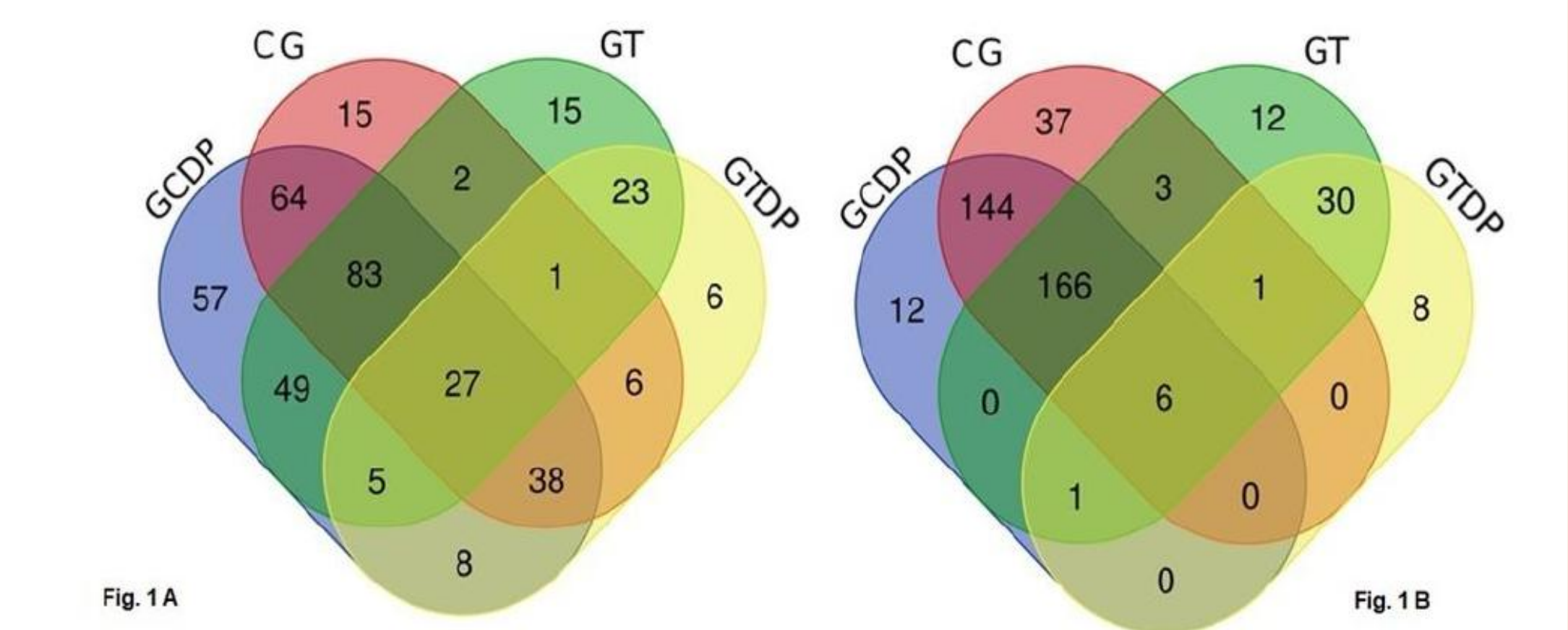
Exclusion criteria: Patients who (i) were pregnant or lactating; (ii) had received a course of periodontal treatment within the last 6 months; (iii) had fever, lower extremity edema, infection, received corticosteroid pulse therapy, uncontrolled hypertension, dyspnea, (iv) total toothless and (v) were not able to provide consent to participate in the study.

61 patients	GT: kidney transplant recipients without periodontal disease (n = 11);	GCDP: normal renal function with periodontal disease (n = 19)
	GTDP: kidney transplant recipients with periodontal disease (n = 12);	CG: normal renal function without periodontal disease (n = 19)

CONCLUSION

- ✓ There was no significant difference between the salivary and urinary metabolites when compared among the four groups;
- ✓ It was possible to differentiate the specific metabolites for each group and the relationships between them;
- ✓ The metabolites expressed in groups with periodontal disease were related to bacterial metabolism and anaerobic cell respiration;
- ✓ The most important metabolite for determining the pattern of PCA performed in saliva and urine samples was ornithine-d7-4 and palmitoleic acid, respectively

RESULTS



PERMANOVA		
	Saliva	Urine
Permutation N:	9999	9999
Total sum of squares:	1.91 × 10 ¹⁹	7.82 × 10 ¹⁹
Within-group sum of squares:	1.84 × 10 ¹⁹	7.23 × 10 ¹⁹
F	0.7176	1.574
P	0.8741	0.103

p < 0.05 indicates the significant difference between groups.

REFERENCES

Gawron K, et al. Metabolomic status of the oral cavity in chronic periodontitis. *In Vivo* 2019; 33 (4): 1165–74.
Huang Y, et al. Mass spectrometry-based metabolomic profiling identifies alterations in salivary redox status and fatty acid metabolism in response to inflammation and oxidative stress in periodontal disease. *Free Radic Biol Med.* 2014; 70: 223–32. PMID: 24607715. DOI: 10.1016/j.freeradbiomed. 2014.02.024.
Xia T, et al. Targeted metabolomic analysis of 33 amino acids and biogenic amines in human urine by ion-pairing HPLC-MS/MS: biomarkers for tacrolimus nephrotoxicity after renal transplantation. *Biomed Chromatogr.* 2018; 32 (7): e4198. doi: 10.1002/bmc.4198.
Zheng L, et al. GC/MS-based urine metabolomics analysis of renal allograft recipients with acute rejection. *J Transl Med.* 2018; 16: 202. <https://doi.org/10.1186/s12967-018-1584-6>.