

Biosensors

B. Tech.

Course No.: EEL 3050

L-T-P [C]: 3-0-2 [4]

Prof. AJAY AGARWAL

ELECTRICAL ENGINEERING

IIT JODHPUR

Lecture 13 dated 5th Sep. 2024

Surface-Enhanced Raman Spectroscopy or Surface-Enhanced Raman Scattering (SERS)

- a surface-sensitive technique that enhances Raman scattering by molecules adsorbed on rough metal surfaces or by nanostructures
- the enhancement factor can be as much as 10^{10} to 10^{11} ,
- the technique may detect single molecules

SERS

- The **exact mechanism** of the enhancement effect of SERS is still a **matter of debate**
- There are **two primary theories** and while their mechanisms differ substantially, distinguishing them experimentally
- the **electromagnetic theory** proposes **the excitation of localized surface plasmons**, while
- the **chemical theory** proposes the **formation of charge-transfer complexes**. It is based on **resonance Raman spectroscopy**, in which the **frequency coincidence (or resonance)** of the **incident photon energy & electron transition** greatly **enhances Raman scattering intensity**.

3. Nano-structure array for *Surface Enhanced Raman Spectroscopy*



Sensors and Actuators A: Physical
Volume 139, Issues 1–2, 12 September 2007, Pages 36–41

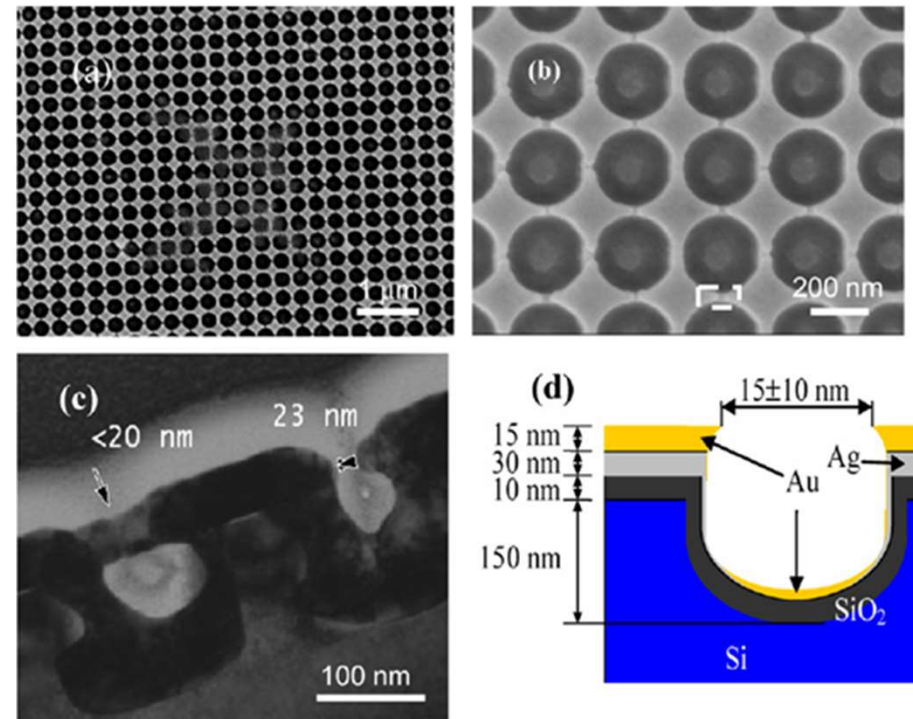


3D arrays of SERS substrate for ultrasensitive molecular detection

R.Z. Tan^a, A. Agarwal^a ✉, N. Balasubramanian^a, D.L. Kwong^a, Y. Jiang^b, E. Widjaja^b, M. Garland^b

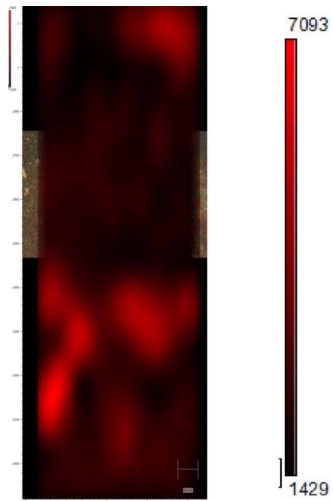
Suitable for trace level detection of:

- Biological warfare,
- Bio-markers,
- Explosives, etc.



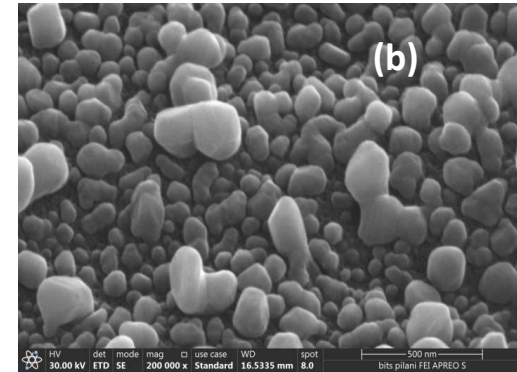
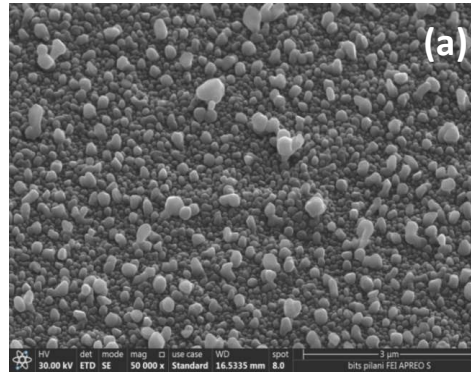
SERS substrates

Bio-sample analysis on SERS substrate

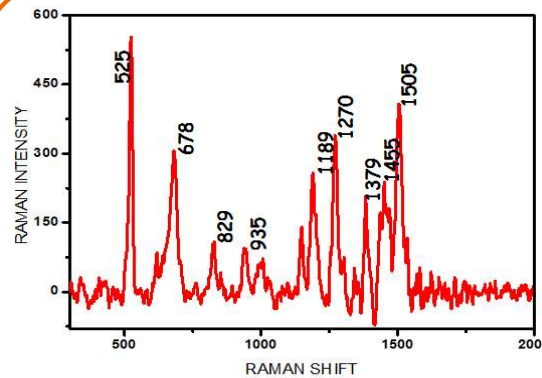


1.5mm x 0.5mm

Raman Mapping of
SERS substrate for
Rhodamine B (10 μ M,
785 nm) at 620 cm^{-1}

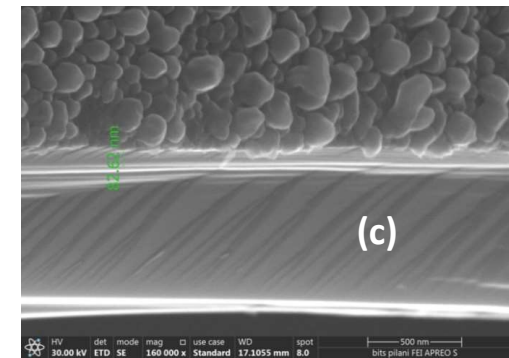


FESEM images of Si SERS substrate surface (a to b)



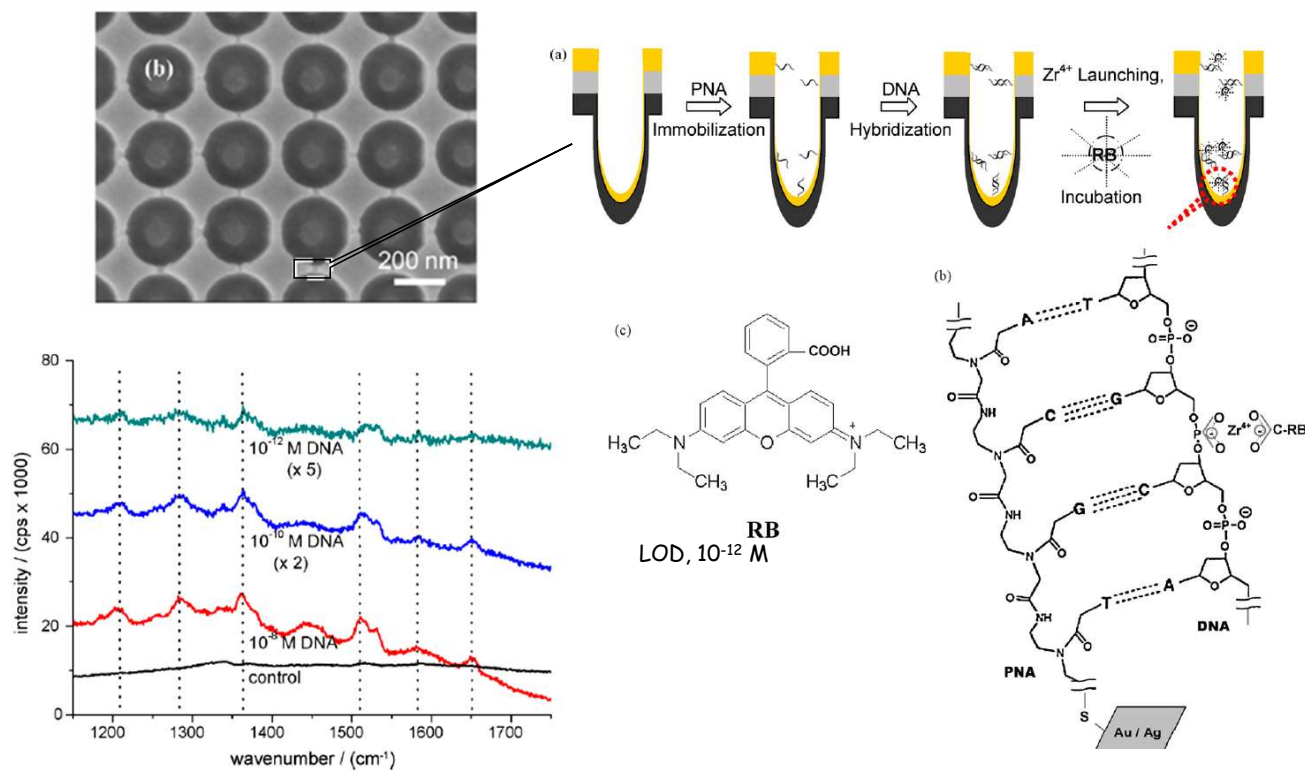
ER-PR-HER-2⁺ TISSUE SAMPLE

Bio-sample analysis of breast
cancer tissue cells



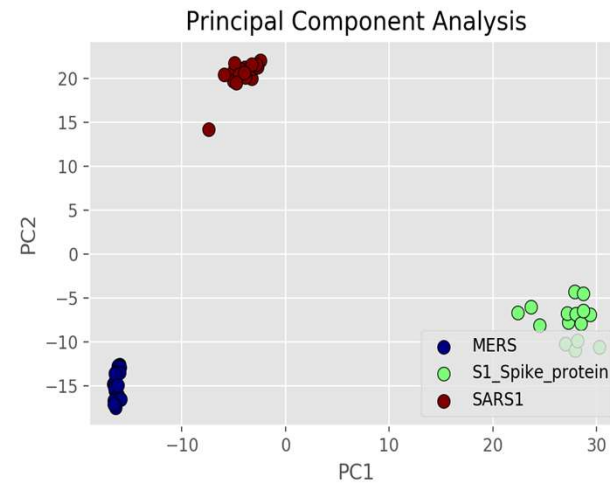
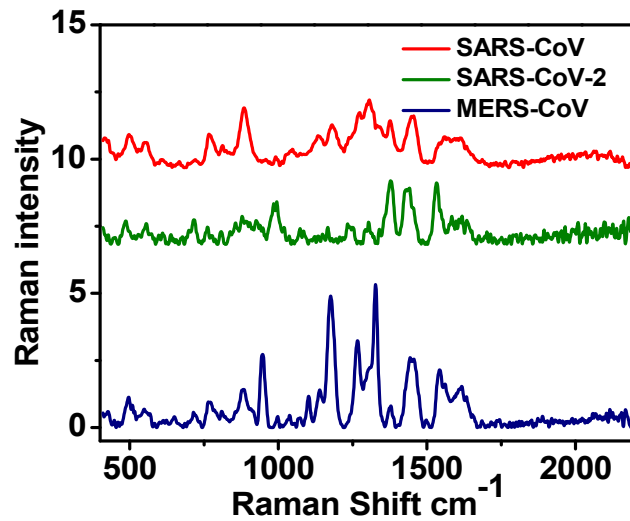
Cross-sectional view

DNA detection

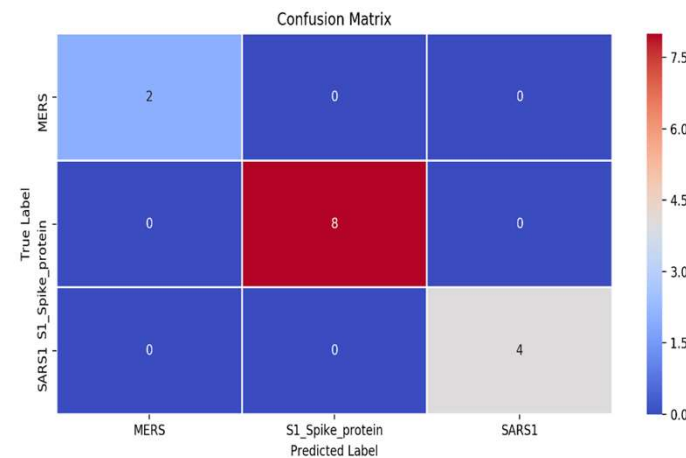


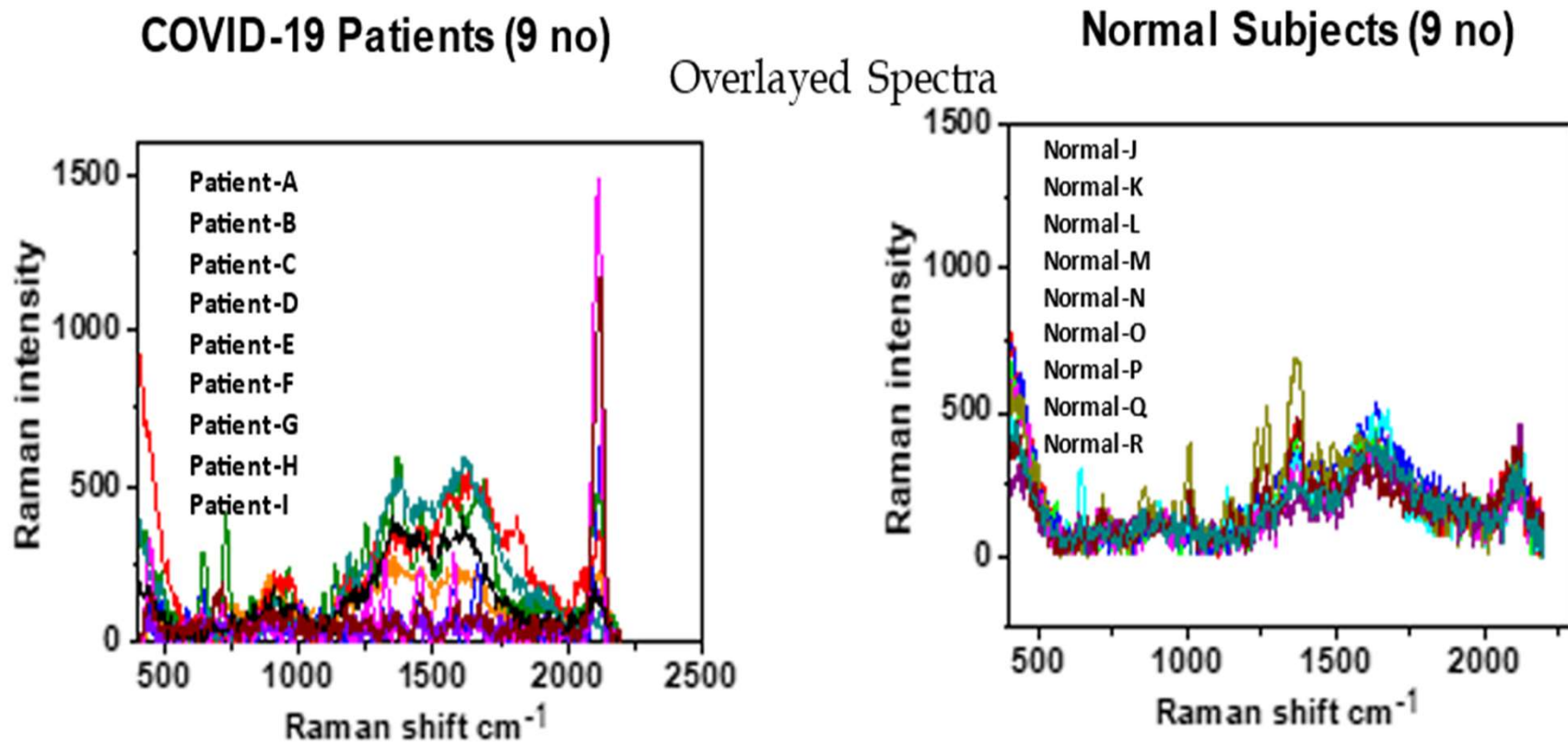
C. Fang, Ajay Agarwal, et al., *Biosensors and Bioelectronics*, 24 (2008) 216–221

Differentiation of Raman fingerprints of three different Corona virus Spike Protein (**SARS-CoV-2**, **SARS-CoV**, **MERS-CoV**) by **PCA & SVM**



Wavenumber cm ⁻¹	Peak assignment
883	CH2 protein assignment
937	Side chain vibration of Proline
1176	C-H bending tyrosine
1327	Amide III



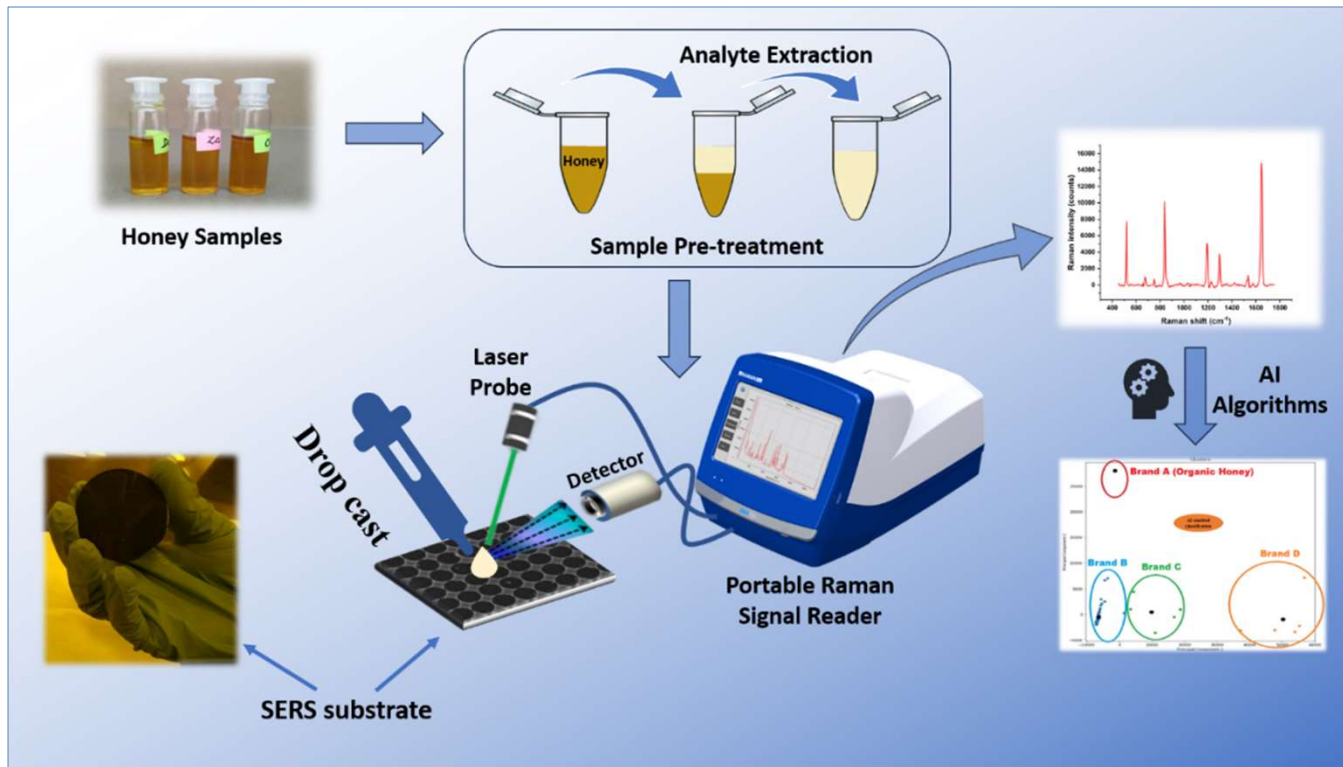


Raman spectra, taken using SERS active substrates, of COVID-19 patients' and normal subjects' saliva

Varsha K, ... , **Ajay Agarwal**, S Singh, Kaustabh K Maiti, A non-invasive ultrasensitive diagnostic ..., Journal of Photochemistry and Photobiology B: Biology, 2022, 112545.

Nanosensor for the Classification of Organic Honey

- Indian Honey market is valued at **INR 2330 Crores** (2022) and expected to reach **INR 3880 Crores** by 2028 (8.4 % CAGR).

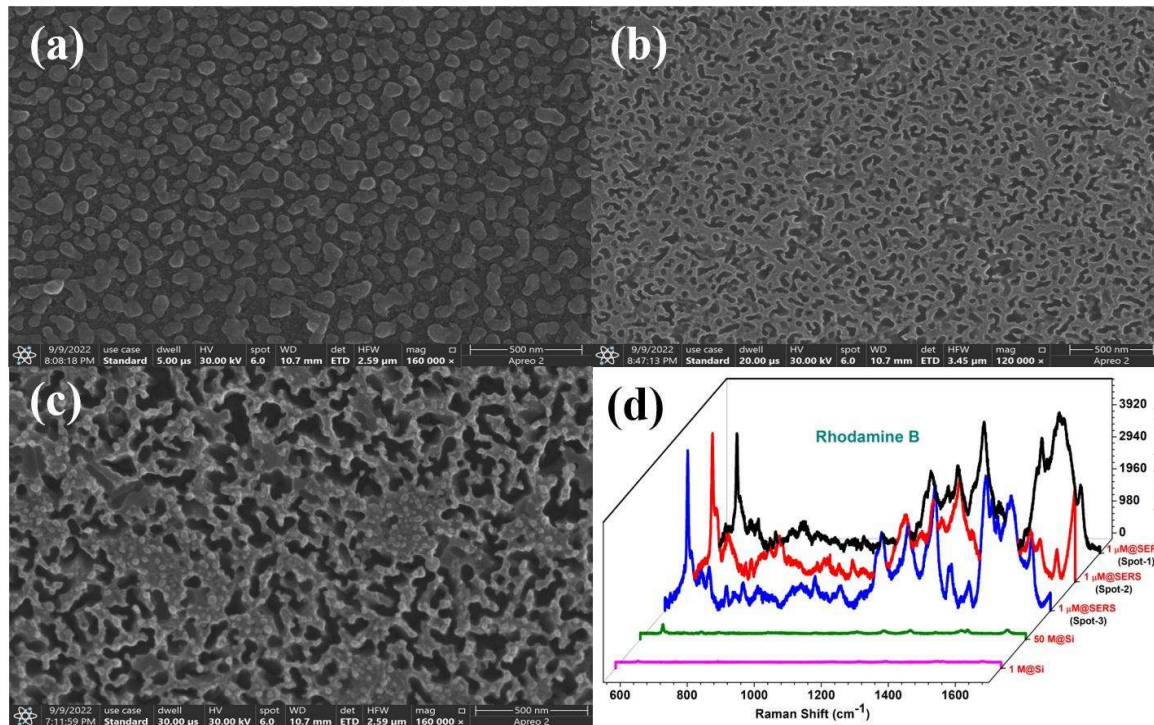


Specifications of nanosensor

- Plasmonic feature size: 20nm – 200nm
- Enhancement factor: 10^6 – 10^9
- Limit of detection: ppm/ ppb
- Target AI algorithms for integration: CNN, SVM or EA

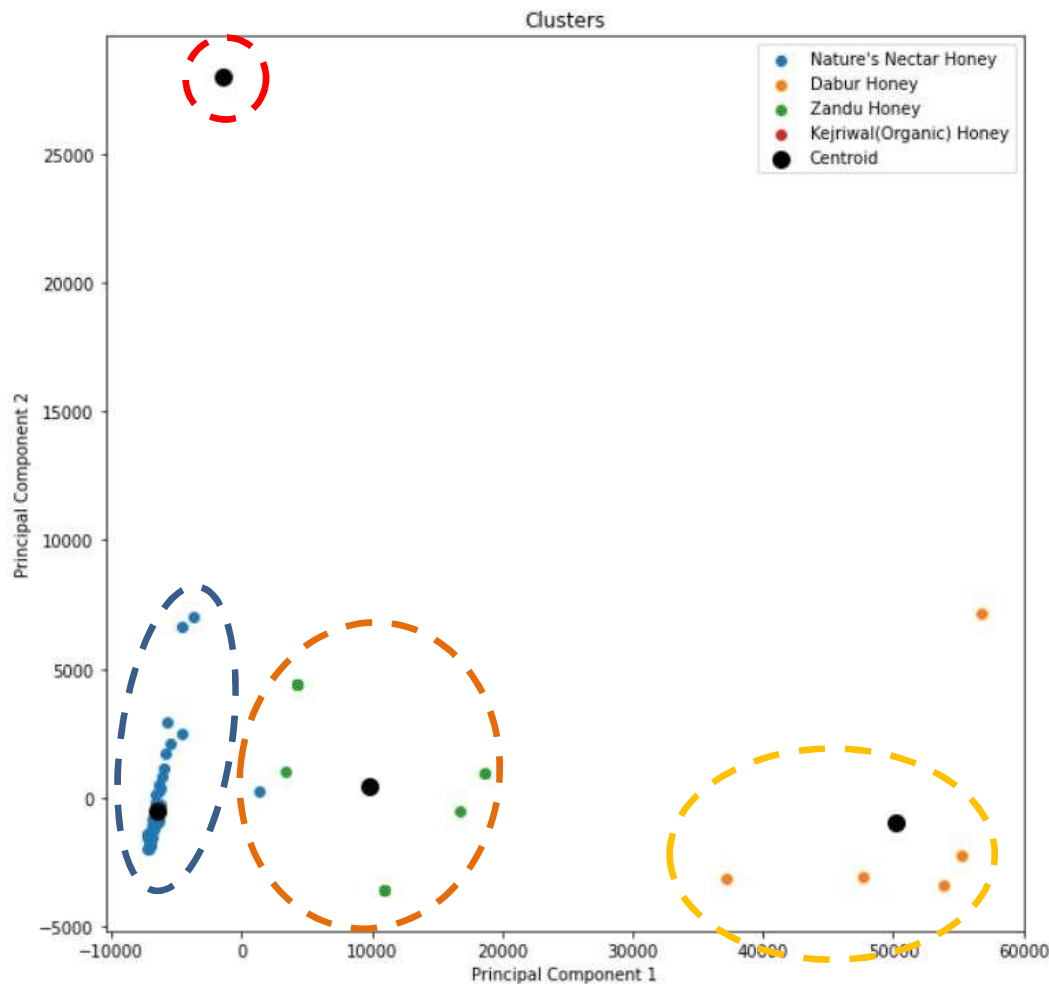
S. Singh, A. Agarwal, et al., "Rapid Detection of Paraquat Pesticide in Honey using SERS-Based Portable Nanosensing Platform," in IEEE Sensors Letters, vol. 7, no. 10, pp. 1-4, Oct. **2023**, Art no. 6006804

Initial results:



- FESEM images of Si SERS substrate after
- (a) Ag NP deposition on planar Si surface
 - (b) porous Si fabrication by chemical etching
 - (c) Ag NP deposition on porous Si
 - (d) SERS behavior of nano-sensor with Rhodamine B (RhB) at 03 spots compared with bulk RhB on planar Si

K-means Algorithm for Honey Classification



- ☐ Asymmetric Partial Least Squares method was used for Spectra smoothing
- ☐ K-means clustering model was used for auto-classification of the four honey types using Raman Spectroscopy
- ☐ The accuracy can be improved by using other deep learning models such as convolutional neural networks etc.

Questions?