Killing 2 birds with one stone: Exploring use of anti-sperm antibody to prevent Cell-associated HIV transmission in addition to pregnancy

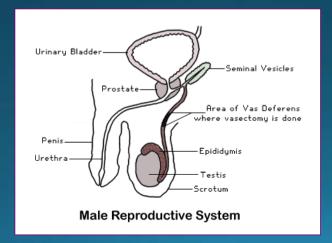
Jai Marathe

Anti-sperm antibody (ASA)

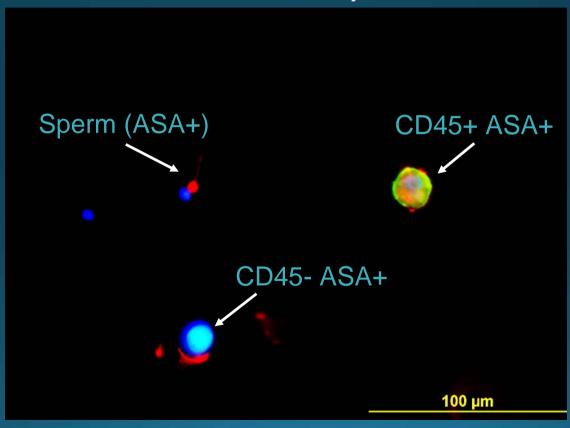
- Discovered in 1980's
- Isolated from blood of an infertile woman
- Agglutinating Antibody directed against CD52g antigen
- CD52g is secreted in male genital tract (Mainly in epididymis, vas deferens and seminal vesicles)

Inserted into the cell-membrane of spermatozoa in the lumen of male

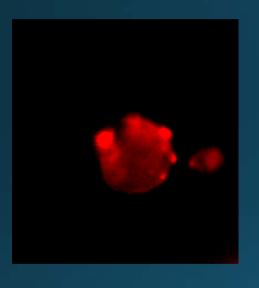
genital tract

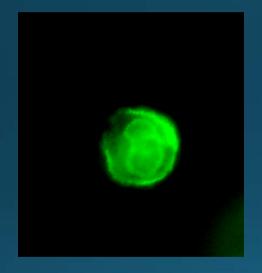


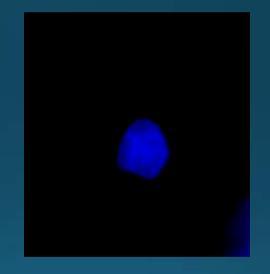
Is CD52g inserted into the cell membrane of seminal leukocytes?

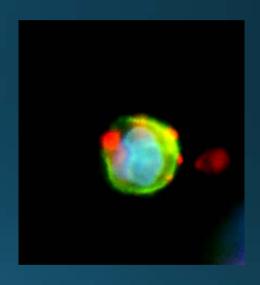


CD45+ ASA+ seminal leukocyte









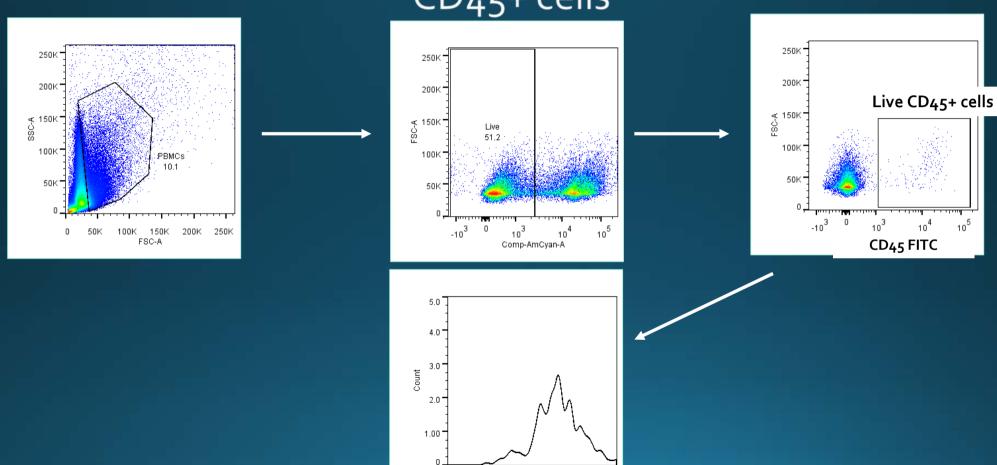
Anti- CD52g (MHS8)

Anti- CD45

DAPI

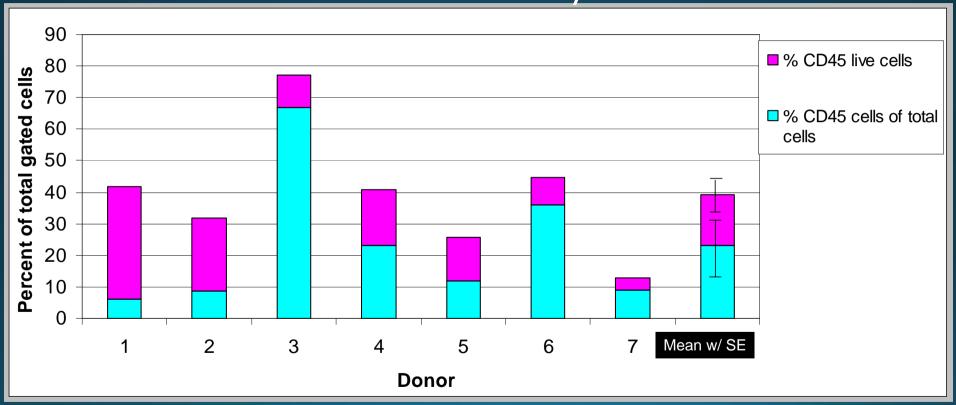
Merge

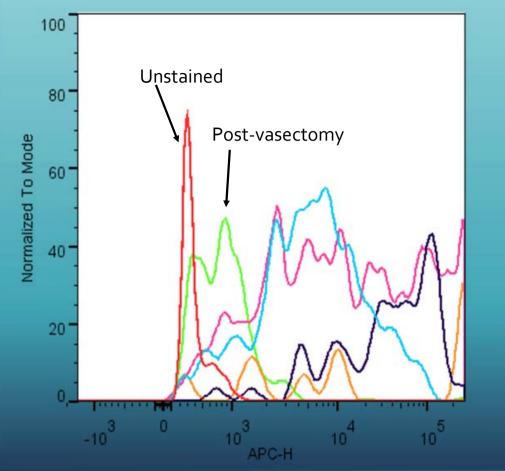
Flow Cytometric Analysis of ASA staining of seminal live CD45+ cells

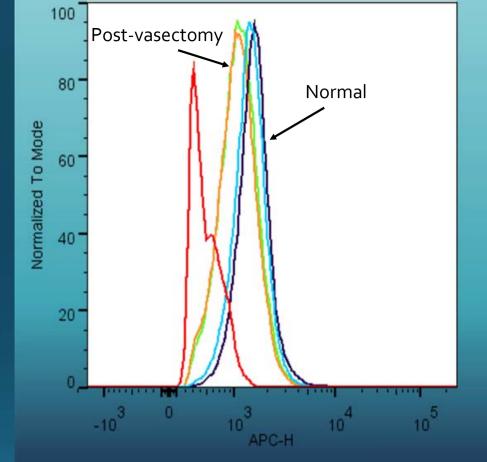


ASA

Interdonor variability in percent live CD₄₅+ seminal leukocytes





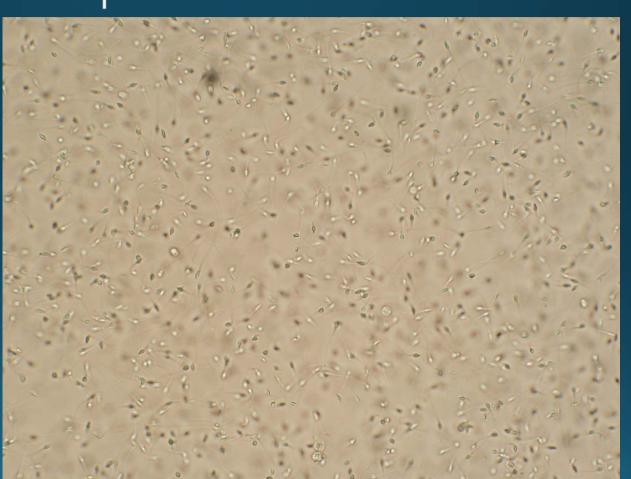


Seminal mononuclear cells isolated on Ficoll gradient stained with CD45 and ASA for flowcytometric analysis

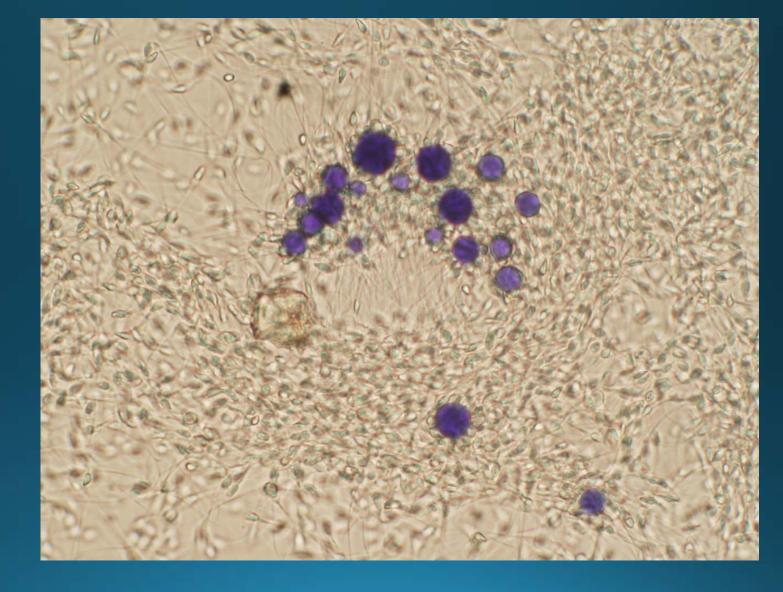
PBMCs incubated for 2 hours with normal or vasectomized seminal plasma stained with CD45 and ASA

Are seminal leukocytes capable of being agglutinated with sperm when treated with ASA

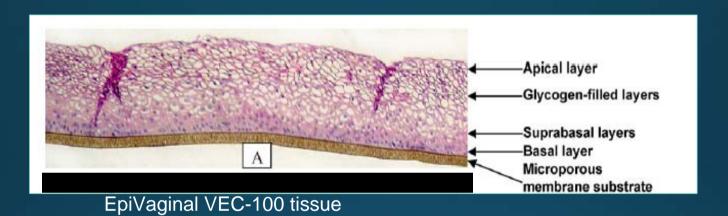
Normal motile sperm



Sperm and live seminal mononuclear cells treated with ASA



Vaginal epithelial model by MatTek



Apical layer

Glycogen-filled layers

Suprabasal layers

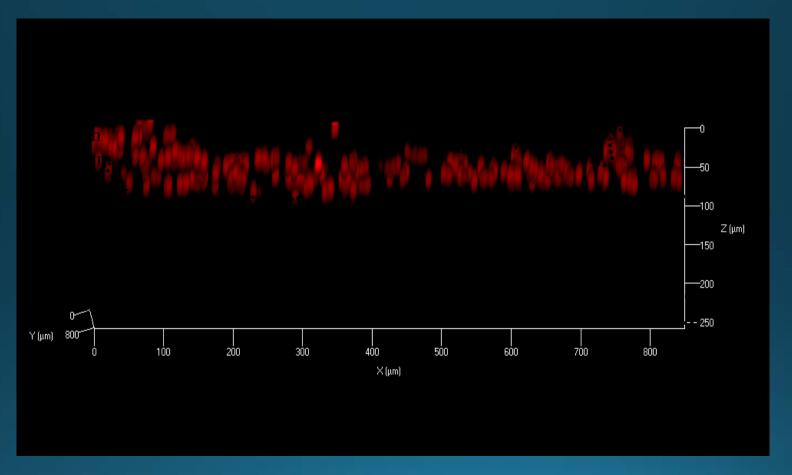
Basal layer

Lamina propria

and other cells

containing fibroblasts

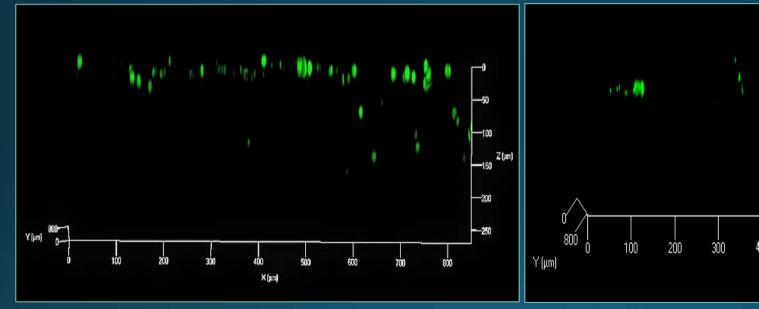
3D Reconstruction: 10um beads on EpiVaginal surface

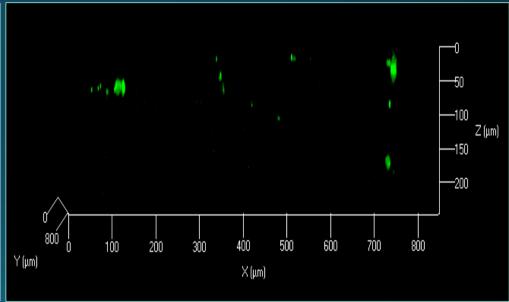


Infiltration studies: Preliminary data

A MDM infiltration in presence of SP

B Seminal leukocyte infiltration in presence of SP





- EpiVaginal tissues were TNF treated for 24 hours prior to infiltration
- Images show cells stained with Live Cell tracker CMFDA

Conclusions

- CD52 g antigen is detected on cell membrane of seminal leukocytes
- >95 % Seminal mononuclear cells were associated with agglutinated spermatozoa on treatment with anti-sperm antibody
- Future studies will determine whether ASA can block mononuclear cell infiltration of tissue

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Thank You!

