nPOD Viral Workgroup Introduction/Overview



Alberto Pugliese







The nPOD Research Model

- Identify synergies, develop project collaborations
- Promote collaboration through focused, coordinated working groups
 - Coordinate study of nPOD cases, throughout the disease spectrum, for integrated studies by multiple investigators
 - Exploit the key unifying element of the shared sample
 - Develop coordinate studies with diversified, complementary approaches and collective input to study design
 - "real-time" data sharing; as activity of these working groups progresses, findings emerge and are shared, informing the group about any changes in strategies and focus on the next questions
 - Web-based connectivity, on line Pathology, LabKey (soon)



Network for Pancreatic Organ
Donors with Diabetes



Key Questions posed by the nPOD Viral Group

- Can enteroviruses or other viruses be identified, isolated?
- Are there replication-defective enteroviruses?
- Is there an acute and/or or chronic (persistent) infection?
- Do viruses infect ß-cells, and/or other pancreatic cell types?
- Could the same viruses detected in pancreas infect immune cells and hence lymphoid tissues? Could this have consequences on immune function?
- Is peripheral blood infection a surrogate marker of viral infection in the pancreas/lymphoid tissues, and a tool to isolate disease-relevant viruses?
- Is there a link between ß-cell proliferation and enterovirus infection?







Diversified Approach to the Questions

<u>Immunostaining</u>

- Validation of antibody reagents
- Reference specimens (human islets, cell lines, mouse tissues)
- Links with insulitis, inflammatory changes (class I, chemokines), regeneration

Immunology

- Study islet antiviral protein responses
- Study virus presence in lymphoid tissues and blood
- Study EVreactive T cell responses
- Viral content in autoreactive T cells

RNA Studies

- In situ hybridization
- RNA-seq (whole pancreas, LCM islets), cDNA sequencing
- Microarrays
- Metagenomics/ computational subtractive analysis
- EV PCR, nested PCR

Protein Studies

 Mass Spec analysis of immunoreactive proteins

Isolating Virus

 Infection of cell lines and mice from infected blood samples, islets, pancreas, supernatants, RNA etc.







nPOD Viral Group Members (in no particular order and growing)

- Sarah Richardson
- Noel Morgan
- Heikki Hyoty
- Maarit Oikarinen
- Nora Chapman
- Matthias Von Herrath
- Ken Coppleters
- Sally Kent
- Antonio Toniolo
- Gun Frisk
- Roberto Gianani
- Nora Chapman

- Dirk Homann
- Suparna Sarkar
- Vincent Plagnol
- David Schneider
- Skip Virgin
- Joe Petrosino
- Richard Lloyd
- John Kaddis
- Martha Campbell-Thompson
- Irina Kusmarsteva
- Emily Montgomery
- Li Zhang
- Amy Wright









nPOD Junior Investigator Award

The Network for Pancreatic Organ Donors with Diabetes acknowledges

Sarah Richardson

for being a champion of collaborative spirit and data sharing and for her dedication to the field of type 1 diabetes research



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