

### **IMP761** webcast slides

Date & Time: March 26, 2019, 7:45 am Australian Eastern Daylight Time

March 25, 2019, 4:45 pm US Eastern Daylight Time

Register: Interested parties can register via a link to the webcast on the Company's website or via the following link: <a href="https://fnn.webex.com/fnn/onstage/g.php?MTID=e3a09aba7876417a76f919285ab7b26bb">https://fnn.webex.com/fnn/onstage/g.php?MTID=e3a09aba7876417a76f919285ab7b26bb</a>

A replay of the webcast will also be available at www.immutep.com from the day after the event.

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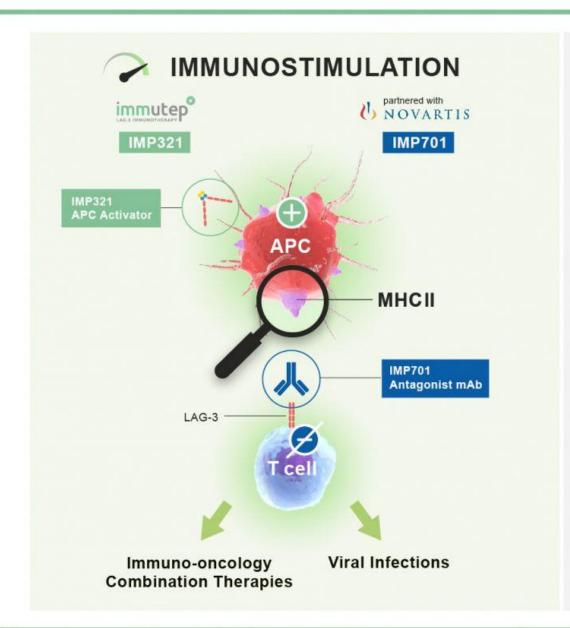
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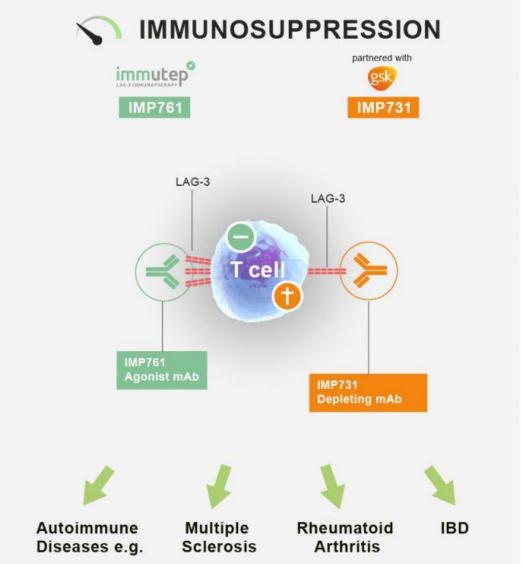
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# Targeting: LAG-3/MHC II May Lead to Multiple Therapeutics in Numerous Indications







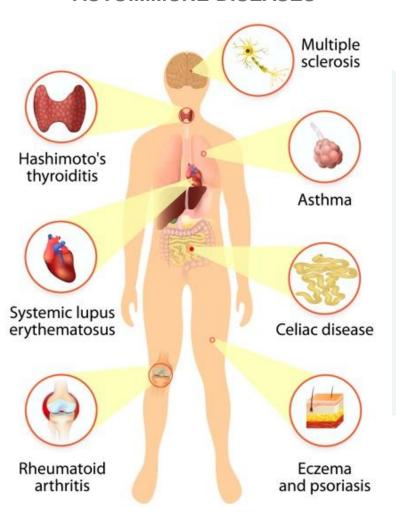


# The LAG-3 Agonist Concept

## Targeting auto-reactive memory T cells in AID with IMP761



#### **AUTOIMMUNE DISEASES**





Treating general inflammation: corticoids, methotrexate, anti-TNF-α, -IL-6, -IL-17, -IL-23 mAbs

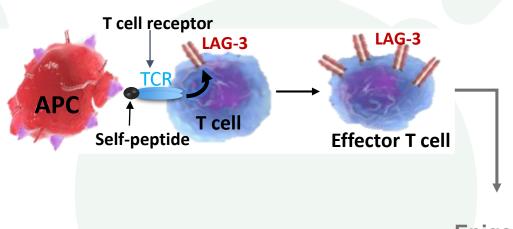


Treating the disease process:
silencing the few auto-immune memory
T cells accumulating at the disease site
with IMP761

## IMP761 works upstream from current therapeutic interventions



Auto-immune memory T cells are chronically stimulated by the same self-peptide, acquiring an 'exhausted' phenotype and expressing LAG-3 which down-modulates specifically TCR signaling. IMP761 increases this physiological down-regulation.



IMP761: blocking the activation of self-reactive memory T cells



#### **Epigenetic reprogramming**

(DNA methylation, histone modifications, miRNAs)



Th1 (e.g. RA, T1D)

Th2 (e.g. allergic asthma)

Th17 (e.g. IBD)

IL-23 (e.g. psoriasis)



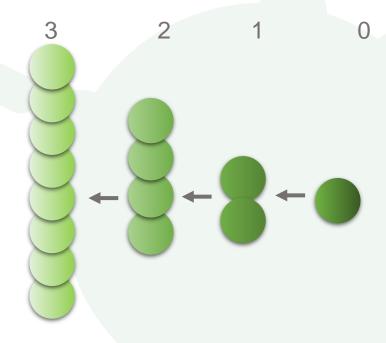
# IMP761: In Vitro Testing

### Inhibition of CD8 T cell proliferation

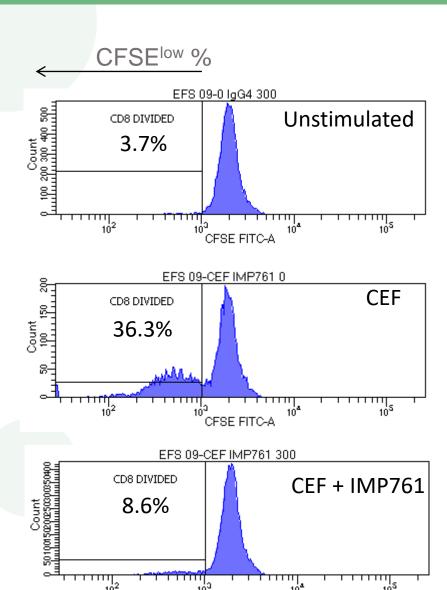


#### CFSE proliferation assay

#### Divisons:



CEF peptide pool CMV – EBV – Influenza Stimulates CD8 T cells in hPBMCs



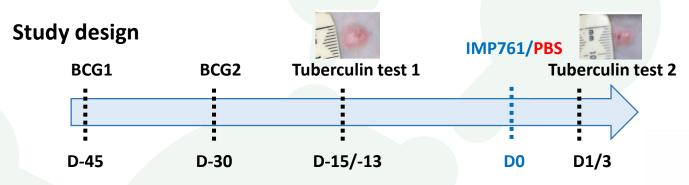
CFSE FITC-A



# IMP761: In Vivo Testing

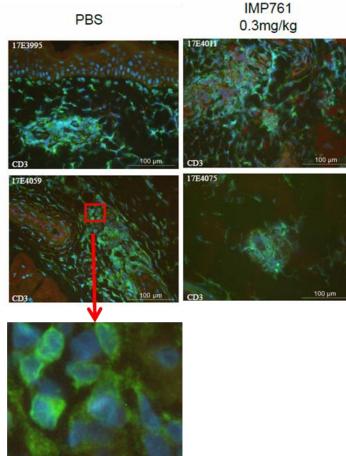
### **Delayed-type hypersensitivity model** in cynomolgus monkey





#### Immunofluorescence staining:

inflammatory T cell infiltration at tuberculin test site before and after IMP761/PBS injection

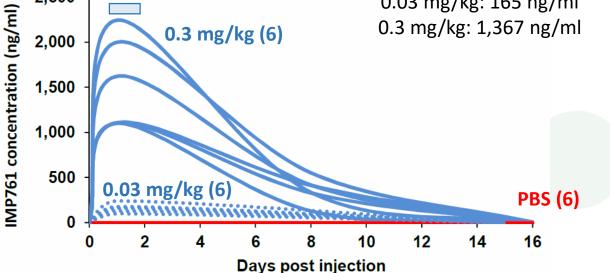


#### **Pharmacokinetics**



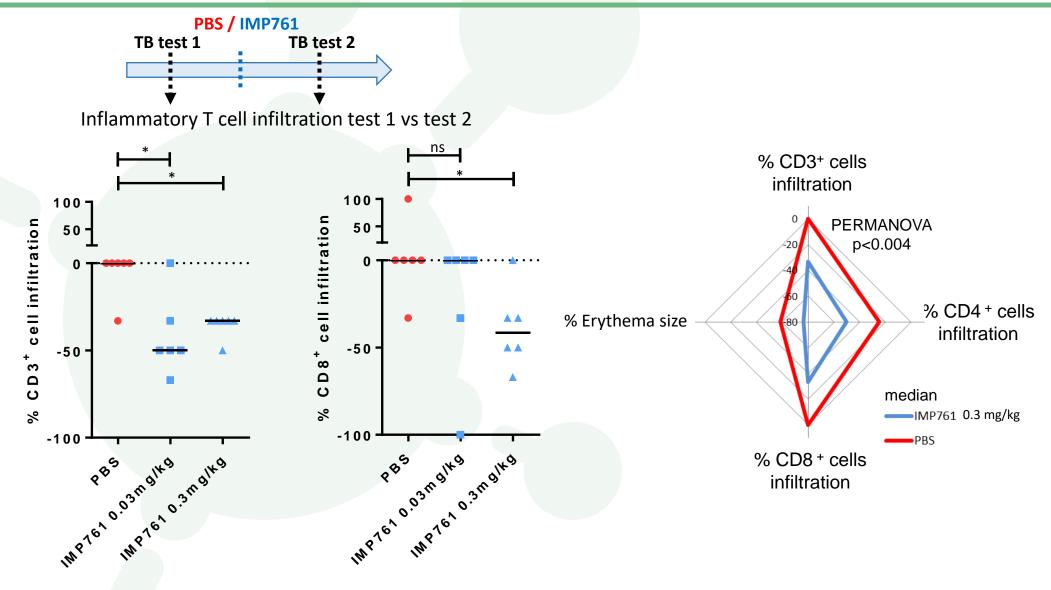
test 2

test 1



## IMP761 inhibits inflammatory T cell infiltration *in vivo*





IMP761 is able to inhibit significantly T cell infiltration of an antigen-specific intradermal reaction

### Conclusion



- The Concept: treating the cause of autoimmune diseases, not just the symptoms
- The Target: the self-peptide specific memory T cells harboring LAG-3
- **The Tool**: an agonistic LAG-3-specific mAb down-modulating self-peptide-induced TCR signaling
- The Evidence (1): in vitro down-modulation of peptide-induced human T cell proliferation and activation
- The Evidence (2): *in vivo* down-modulation of peptide-induced T cell infiltration/inflammation at the tissue site in a NHP model
- The Status: Cell line development ongoing and GMP manufacturing preparations underway in order to progress to clinical development.



# Thank you!