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## Advancing Personalized Healthcare with High-Performance Cloud Computing for the Living Heart Project

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#### Case Study: The Living Heart Project

Studying Drug-induced Arrhythmias of a Human Heart with Abaqus in the Cloud







### The Living Heart Project



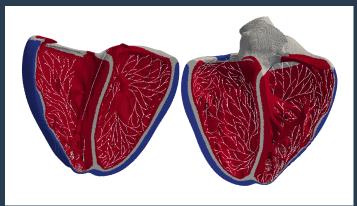
- Members: Leading cardiovascular researchers, educators, medical device developers, regulatory agencies, and practicing cardiologists
- Shared mission to develop and validate highly accurate personalized <u>digital human heart models</u> (DHHM).
- Living Heart Models establish a unified foundation for cardiovascular in silico medicine
- Models serve as a common technology base for education and training, medical device design, testing, clinical diagnosis and regulatory science
- Rapidly translating current and future cutting-edge innovations directly into **improved patient care**.

#### Arrhythmia affects millions of people

- In Europe and North America, atrial fibrillation affects about 2% to 3% of the population (2014)
- Atrial fibrillation and atrial flutter resulted in 112,000 deaths in 2013, up from 29,000 in 1990
- Sudden cardiac death is the cause of about half of deaths due to cardiovascular disease or about 15% of all deaths globally
- About 80% of sudden cardiac death is the result of ventricular arrhythmias

#### Multiscale model of cardiac electrophysiology

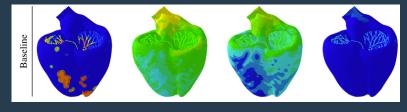
- Bi-ventricular anatomy based on healthy human
- Finite element model with 7,500,000 nodes
- 250,000,000 internal variables, updated/stored within each sim step
- 1,000,000 time steps
- State of the art representation of cellular dynamics
- 3 different cell types in the ventricular wall
- High fidelity model of the Purkinje network

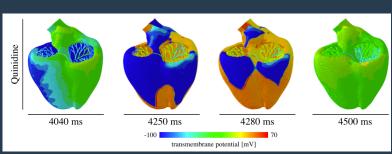


### Applying drugs to the living heart

- Torsades de Points is a dangerous type cardiac arrhythmia. Electrical waves in the heart turn chaotic.
- Produced as side effect of drugs, but assessing this risk for new compounds is expensive and can take a long time
- With this model, we can predict overall response the heart just by measuring the effect of a drug in a single cell
- Example: we applied Quinidine to the model, which presents high risk of Torsades de Points
- Our model spontaneously develops Torsades de Points when high risk drugs are applied
- We envision this model will help researchers, regulatory agencies, and pharmaceutical companies to accelerate drug development and create effective and safe drugs for patients.

#### regular rhythm





Torsades de Points

High Performance Computing as a Service (HPCaaS)

Partners: Advania / HPE / Intel / Dassault / UberCloud

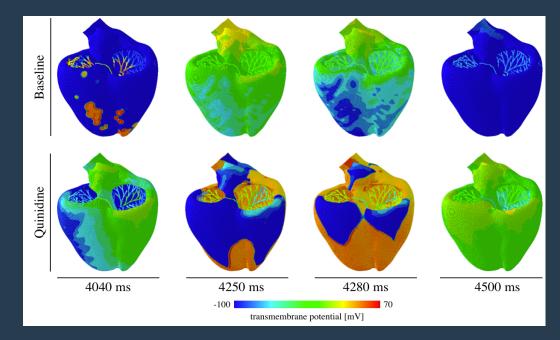
- Advania's HPC as a Service (HPCaaS) hardware configuration
- Built upon 100 HPE ProLiant servers XL230 Gen9
- Each with 2 Intel Broadwell E5-2683 v4 Intel OmniPath interconnect
- UberCloud HPC software **containers** hosting Stanford's workflow
- Dassault Systèmes SIMULIA Abaqus for structure and advanced electro-physiological interaction

HPCaaS Environment and Simulations Advania / HPE / Intel / Dassault / UberCloud

Goal: create a biventricular finite element model for **Stanford** to study drug-induced arrhythmias of a human heart.

- Hundreds of cloud HPC hours on different Advania configurations
- LHP model scaled well up to 240 compute cores
- 42 simulations each 40 hours on 5-node (160-core) subsystem
- Study: identifying drugs causing arrhythmias
- Applying drugs by blocking different ionic currents in cellular model, replicating what has been observed before in cellular experiments
- For each case, we let the heart beat naturally and see if the arrhythmia is developing

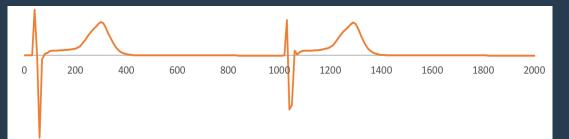
#### **Simulation Results**



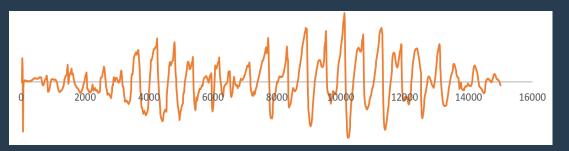
Evolution of electrical activity: After application of Quinidine, the electrical propagation turns chaotic, showing the high risk of Quinidine to produce arrhythmias.

#### Simulation Results

Electrocardiogram (ECG) without and with the drug Sotalol



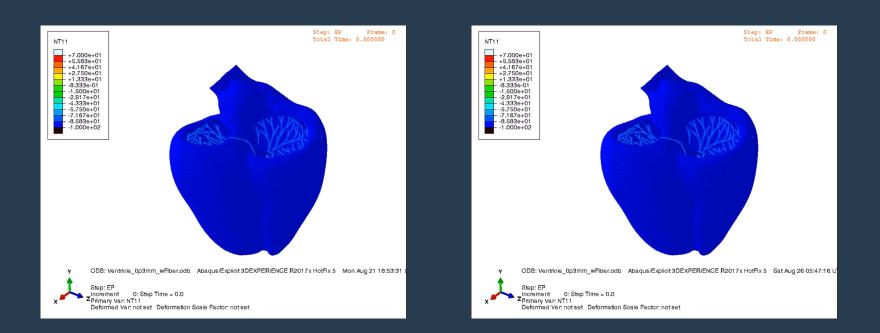
#### ECG tracing for healthy, baseline case



Arrhythmic development after applying the drug Sotalol. The ECG demonstrates that the arrhythmia is of Torsades de Pointes type.

Note: These are simulation results !

#### Videos of healthy case versus drug-induced case



Application of the drug Quinidine (right) where we observe Torsades de Points arrhythmia The two videos can be obtained from wolfgang.Gentzsch@TheUberCloud.com

#### Take Aways

**Hewlett Packard** Enterprise

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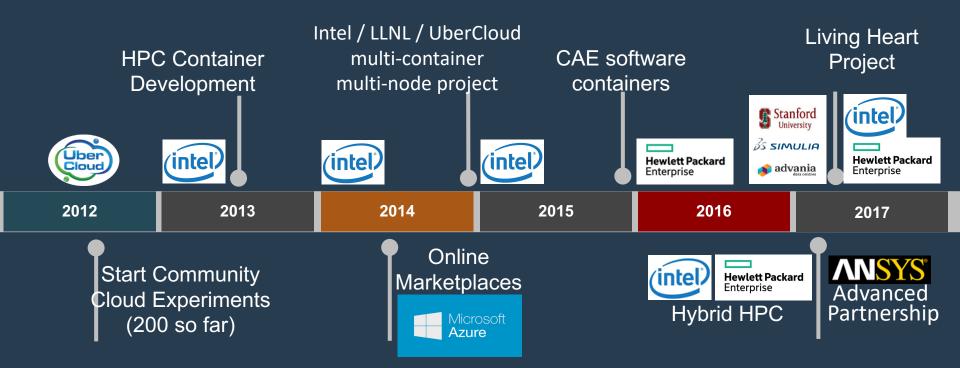




- UberCloud is part of HPE's HPCaaS <u>"Hybrid</u> <u>HPC</u>"
  - HPC <u>Containers</u> give us a way to solve software management problems without performance issues
  - Able to manage and run the most complex engineering <u>workflows</u>
- Providing <u>SaaS-like</u> user experience and desktop level ease of use



### Some Background about the Intel - HPE - UberCloud Partnership 2012 - 2017

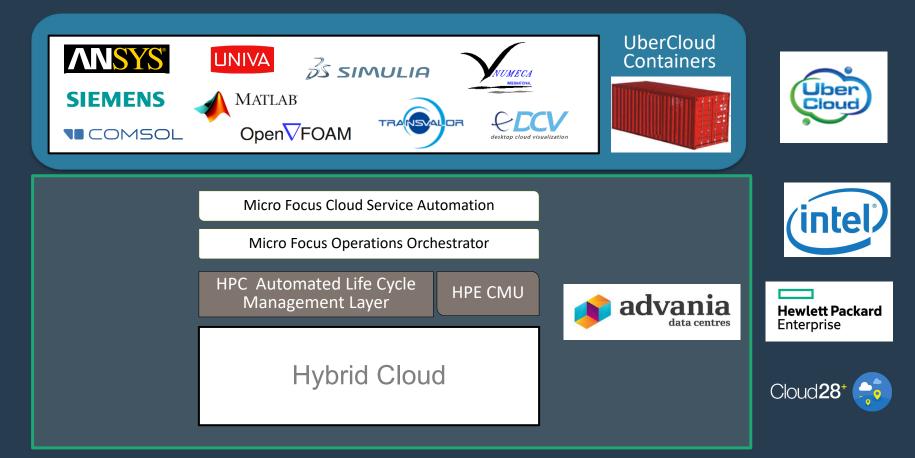


#### Abaqus in UberCloud Container



- Based on Docker, enhanced for engineering & scientific app software
- Application software is pre-installed, configured, tested by UberCloud and Stanford
- Includes **all tools** an engineer needs such as MPI and remote visualization
- Running in the Advania Data Centers
  Cloud

#### LHP Simulations ran on HPE Hybrid HPC Stack



#### Fully Secure Environment on Advania, HPE, UberCloud

HTTPS/VPN Access\*

**OS Firewall** 

**OS PKI Login** 

Network & Storage Segmentation

Dedicated Servers in High Security Data Centers Connection to our servers are protected by strong **encryption** techniques such as HTTPS and VPN (HTTPS/VPN access and Disk Encryption are optional)

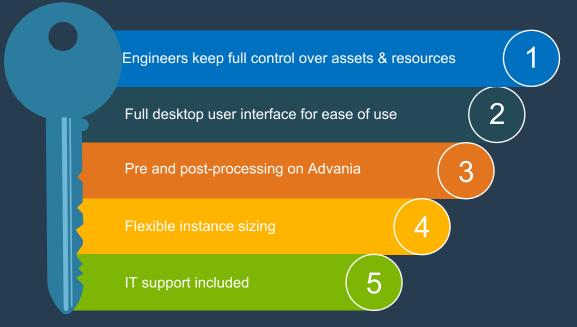
Servers are protected with **firewalls**. Only necessary ports are turned on to reduce attacks

Admin access is protected by **Public Key Encryption** (vs passwords, which can be guessed)

Network and Storage tiers are **segmented** automatically. Each customer gets a private HPE CMU, a private IP range, and a private storage volume

**Single tenant**, bare metal servers (not shared between customers). These servers reside in professionally managed, highly secured data centers.

#### Summary of Key Benefits of Using HPC Containers UberCloud: Looks & works like your desktop only much faster



#### **Usability – Flexibility – Performance**

- Desktop-like user experience to eliminate training needs & provide ease of use
- All software fully installed & ready to use, pre/post processing, meshing, solvers, MPI
- Instance & cluster sizes are flexible, scales
  up/down based on analysis requirements
- Shared storage sized based on needs
- GPU, RDMA, InfiniBand & SSD supported out of the box
- No new cloud platform, no need to learn anything new – feel 'home' immediately

#### Finally, Big Thanks to HPE, Intel, HPCwire And to the HPC User Forum Steering Committee for all the Awards





# Thank You

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