

# ***Project Milestone 1***

**Project Name:** *Biochemical Plate, Assay,  
and Result Management  
system*

**Team:** *West-East99 (We99)*

**Date:** *23<sup>rd</sup> Feb 2015*

**Customer:** *Peter Henstock*

CSCIE-99  
PROJECT



## Contents

<b>1. DOCUMENT ADMINISTRATION.....</b>	<b>3</b>
1.1 DOCUMENT CHANGE HISTORY .....	3
1.2 DOCUMENT CONTENT OWNERS .....	3
1.3 DOCUMENT REVIEWERS .....	4
1.4 DOCUMENT APPROVERS.....	4
<b>2. PROJECT OVERVIEW .....</b>	<b>5</b>
2.1 PROJECT VISION .....	5
2.2 CURRENT PROCESS PROBLEM BEING SOLVED .....	5
<b>3. ASSUMPTIONS AND CONSTRAINTS.....</b>	<b>6</b>
3.1 ASSUMPTIONS .....	6
3.2 CONSTRAINTS.....	6
<b>4. SCOPE.....</b>	<b>7</b>
4.1 IN SCOPE .....	7
4.2 OUT OF SCOPE .....	7
<b>5. PROPOSED FUNCTIONALITY.....</b>	<b>8</b>
5.1 INTRODUCTION.....	8
5.2 OVERVIEW .....	8
<b>6. BUSINESS (FUNCTIONAL) REQUIREMENTS .....</b>	<b>9</b>
6.1 SECURITY FUNCTIONAL REQUIREMENTS.....	9
6.1.1 Security Overview.....	9
6.1.2 Sample Screens / Story Board .....	9
6.1.3 Open Issues / Questions Security Section .....	11
6.2 PLATE MANAGEMENT REQUIREMENTS .....	12
6.2.1 Plate Management Overview .....	12
6.2.2 Sample Screens / Story Board .....	14
6.2.3 Open Issues / Questions Plate Management.....	21
6.3 EXPERIMENT MANAGEMENT REQUIREMENTS .....	22
6.3.1 Overview.....	22

6.3.2	Sample Screens / Story Board .....	22
6.3.3	Open Issues / Questions Experiment Management .....	23
<b>6.4</b>	<b>RESULTS ANALYSIS REQUIREMENTS .....</b>	<b>24</b>
6.4.1	Overview .....	24
6.4.2	Sample Screens / Story Boards .....	25
6.4.3	Open Issues / Questions Results Analysis .....	31
<b>6.5</b>	<b>SPECIAL FEATURE PROPOSALS .....</b>	<b>32</b>
6.5.1	Overview .....	<b>Error! Bookmark not defined.</b>
6.5.2	Interactive Visualization .....	<b>Error! Bookmark not defined.</b>
6.5.3	Audit Trail .....	32
6.5.4	Four Eyes Protocol .....	<b>Error! Bookmark not defined.</b>
6.5.5	Flexible Plate Template Editor .....	<b>Error! Bookmark not defined.</b>
6.5.6	Plate Sharing .....	<b>Error! Bookmark not defined.</b>
6.5.7	Support Machine / Equipment File parsers .....	<b>Error! Bookmark not defined.</b>
6.5.8	Open Issues / Questions Special Features .....	33
<b>7.</b>	<b>HIGH LEVEL DESIGN .....</b>	<b>34</b>
<b>7.1</b>	<b>ARCHITECTURE .....</b>	<b>34</b>
7.1.1	Introduction .....	34
7.1.2	Model .....	35
7.1.3	Controller .....	35
7.1.4	View .....	35
7.1.5	Components .....	35
7.1.6	Database .....	36
<b>8.</b>	<b>PROJECT PLANNING AND ESTIMATING .....</b>	<b>37</b>
<b>8.1</b>	<b>ITERATIONS .....</b>	<b>37</b>
8.1.1	Overview .....	37
8.1.2	Iteration Calendar .....	37
<b>8.2</b>	<b>STORIES AND ESTIMATES .....</b>	<b>38</b>
<b>8.3</b>	<b>MAIN DELIVERABLES .....</b>	<b>38</b>
<b>9.</b>	<b>PROJECT RISKS .....</b>	<b>39</b>
<b>9.1</b>	<b>RISKS IDENTIFIED .....</b>	<b>39</b>
<b>10.</b>	<b>APPENDIX STORY BACKLOG MARCH 2ND .....</b>	<b>40</b>

## 1. DOCUMENT ADMINISTRATION

### 1.1 DOCUMENT CHANGE HISTORY

DATE	AUTHOR	VERSION	DESCRIPTION OF CHANGE
23 <sup>rd</sup> February 2015	Alan Orcharton	1.0	Initial Document
24 <sup>th</sup> February 2015	Sean Sinnott	1.1	Added information to section 5.4 (results import) – and some questions related to the selection of points for linear regressions.
25 <sup>th</sup> February 2015	Alan Orcharton	1.2	Merge comments from Mark and Alex
1 <sup>st</sup> March 2015	Alex Zaman	2.2	Updated Plate Editor Text and mockups.
1 <sup>st</sup> March 2015	Alan Orcharton	2.3	Updated Project Overview sections.
3rd March 2015	Alan Orcharton	2.4	Updated Plate Editor Text and mockups.
3rd March 2015	Alan Orcharton	2.5	Updated Experiment Mgr and Architecture. Also added stories and estimates.
3rd March 2015	Alan Orcharton merged Sean Sinnott changes	2.6	Updated analysis section.

### 1.2 DOCUMENT CONTENT OWNERS

ROLE	NAME
Author	Alan Orcharton
Contributor	Alexander Zaman
Contributor	Tim Stefanski
Contributor	Mark Ford
Contributor	Sean Sinnott

### 1.3 DOCUMENT REVIEWERS

NAME	ROLE
Peter Henstock	Customer
Robert Zupko	Consultant
Alexander Zaman	We99 Front End Developer
Tim Stefanski	We99 Front End Developer
Sean Sinnott	We99 Front End Developer
Mark Ford	We99 Back End Developer
Alan Orcharton	We99 Back End Developer

### 1.4 DOCUMENT APPROVERS

NAME	ROLE	SIGNATURE / ELECTRONIC APPROVAL	SIGN-OFF DATE
Peter Henstock	Customer		

## 2. PROJECT OVERVIEW

### 2.1 PROJECT VISION

Our overall vision is to create a centralized web application that will provide tools to assist scientists in managing experiment plate sets and with the analysis of experimental results.

The team focus is providing outstanding tools to assist in analyzing the dose response characteristics potential drugs.

We will provide beautiful interactive visualization tools that will assist the scientist in:

- Creating and managing experimental plate sets for dose response experiments
- Performing quality control checks on plate results and plate controls
- Performing Interactive analysis of dose response results.

The team will store the experimental results and the analysis of the results so that the historic experimental analysis can be reviewed or reproduced at any time.

Scientists will be able to save the results for further analysis or publish their analysis, making it available to all other users.

### 2.2 CURRENT PROCESS PROBLEM BEING SOLVED

New drug development is a time consuming and expensive process. In the early stages of drug discovery millions of compounds are screened and tested on the road to finding a few candidates for further exploratory development.

High throughput screening is a method scientists' use in the early stages of drug discovery to test millions of potential drugs. The method allows scientist to quickly identify candidates. While the process is highly automated, a disparate selection software of software tools are often used to manage plate sets and analyze results. Commercial software to manage the process is often extremely expensive

The We99 team is proposing releasing an integrated suite of open source web based tools. The team believes that the tools will enable scientists to be more organized and efficient when screening and analyzing potential drugs. Time savings will allow scientist to more effectively eliminate compounds earlier in the discovery cycle potentially saving companies millions of dollars.



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## **3. ASSUMPTIONS AND CONSTRAINTS**

### **3.1 ASSUMPTIONS**

### **3.2 CONSTRAINTS**

- Software Solution must be free. All source code for the solution is to be made available to the open source community. Any part of the system may be openly used or further developed.





## **4. SCOPE**

### **4.1 IN SCOPE**

### **4.2 OUT OF SCOPE**

Interfacing directly with readers of any other experimental hardware.



## 5. PROPOSED FUNCTIONALITY

### 5.1 INTRODUCTION

The purpose of this document is to assist the We99 project team and the customer gain a common understanding of the high level functionality of the Biomedical Plate, Assay and Result Management system. The requirements have been documented in storyboard format. A short description of the functionality is included with some mockup screen shots. This format is intended to provide a good overview of the system from a users point of view.

### 5.2 OVERVIEW

We propose a web-based software solution to facilitate the creation of Biochemical plate specifications, the management of experiments (assays), and the analysis of results.

We have broken the requirements into sections to facilitate the review and analysis of the functional requirements as follows:

#### **Security**

Requirements associated with logging into the system and defining user roles within the system.

#### **Plate Management**

The definition, creation, and storage of biochemical plates and plate sets to be used in experiments

#### **Experiment Management**

The definition, creation, and storage of experiments conducted.

#### **Results Analysis**

Functionality associated with loading, analyzing, reviewing, and storing experiment results.



## 6. BUSINESS (FUNCTIONAL) REQUIREMENTS

### 6.1 SECURITY FUNCTIONAL REQUIREMENTS

#### 6.1.1 Security Overview

We propose that all users of the system must have

- An email address – for use as a unique identifier and during the login process.
- An account – to logon to the system
- A User role – Administrator or Scientist

Nearly all of the operations in the system require a specific permission granted to the user in order to access the feature. These permissions are grouped into Roles and a User is assigned a single Role. This approach allows for easy expansion of the system to include user defined roles or customization of the existing roles.

A user with the Administrator role has access to create and remove user accounts.

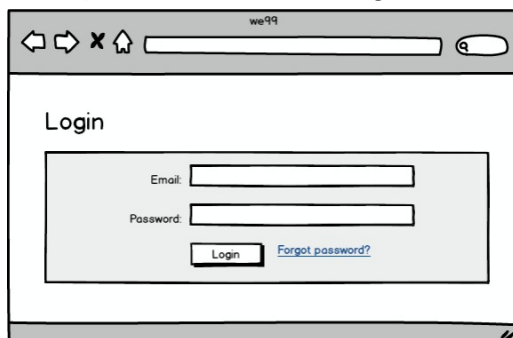
We propose requiring the following information for a user account:

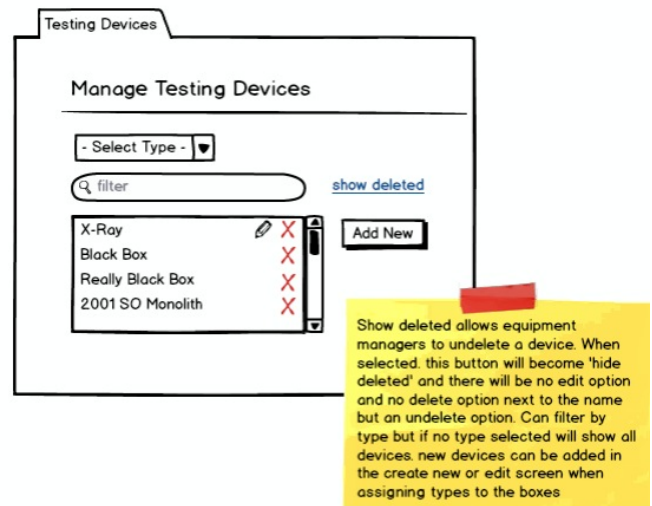
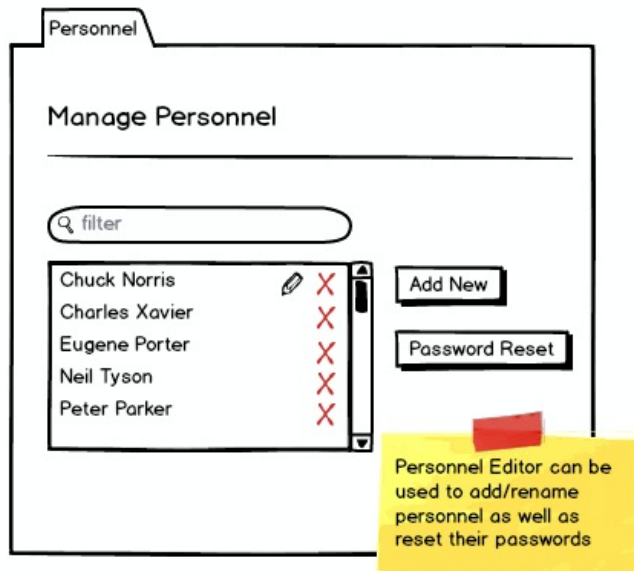
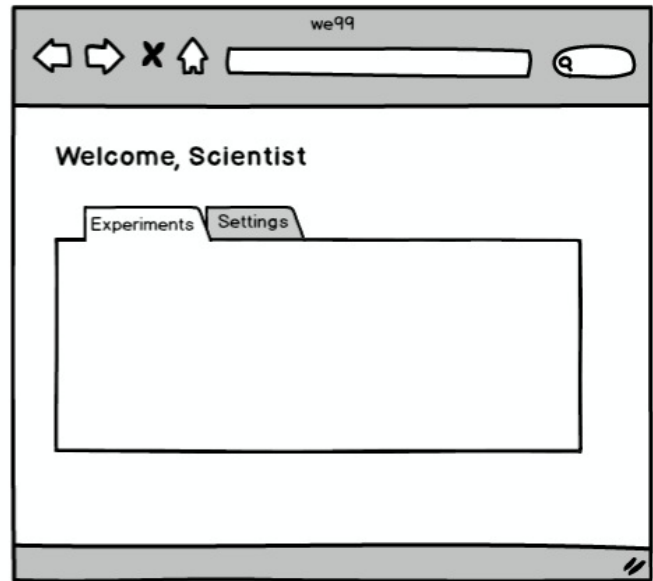
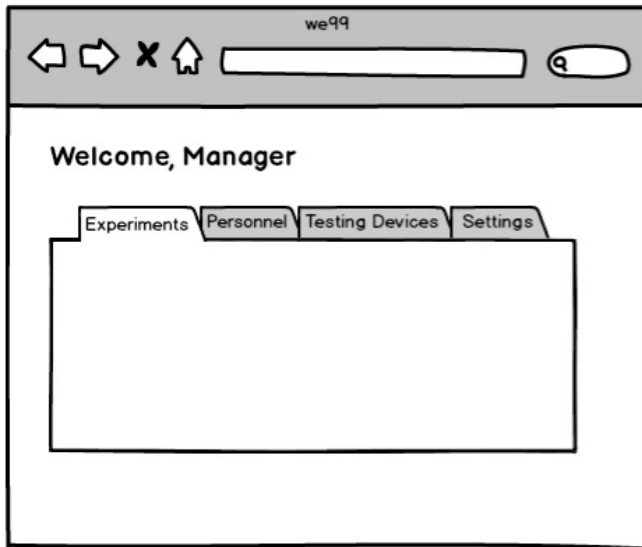
- **Email address** – used as the username. Email addresses help ensure unique usernames. The email address may be used in future for sending notification to the user.
- **First name / Last name** – used to identify the user in a more friendly form.
- **Password** – a user supplied password for authentication.

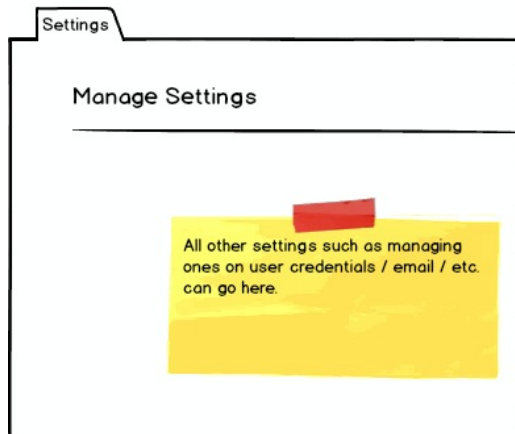
Users with an Administrator role have access to create and remove user accounts and to reset user passwords.

Administrative users also have the ability to add data on test equipment used in experiments. This information can be used to help identify defective equipment and functionality can be expanded to incorporate file parsers to assist the user in loading results produced from a machine that may be in a non-standard format (see special features).

#### 6.1.2 Sample Screens / Story Board







### 6.1.3 Open Issues / Questions Security Section

#	DESCRIPTION	NOTES
1	We can possible use Testing devices to manage import export formats of files associated with the device	
3	Organization Units – Is there a need to allow organizational groupings to manage accounts in large corporate environments	Is a nice idea for large organizations



## 6.2 PLATE MANAGEMENT REQUIREMENTS

### 6.2.1 Plate Management Overview

Plate Management covers creating a specification of the contents of microtitre plate sets to be used in experiments.

The plate management service offers a central place for the user to manage templates and create plate sets for their experiment. The experiment manager offers an overview of existing plate sets and their status. It also offers quick access to create new plate sets.

#### Plate Templates

Plate sets are created using plate templates. Plate templates contain labels that act as place holders for how compounds and controls should be arranged on plates. The process of creating plate templates is covered in detail in the storyboards below.

Plate templates are stored in the system and can be searched based on several criteria.

- Number of positive
- Number of negative controls

#### Plate Details

Plates are identified with a plateID (a.k.a barcode). Plates are assumed to be rectangular and can be created in any rectangular dimension. The dimension is specified using the number of rows and the number of columns. The product of rows x columns defines the number of wells that a plate possesses. Wells themselves are identified by the plate to which they belong and their row and column coordinates.

Plates are made ready for experiment by filling its wells with a specific concentration of a compound.

The concentration of a compound is specified via a number and a unit of measure (e.g., 20 uM).

#### Creating New Plate Sets

New plates can be created using the 'Add New' button from the experiment manager. This leads to a plate set creation wizard. The user uses this wizard to provide relevant information such as:

- The name of the plate set
- Plate type (make / model)
- # of plates and their IDs
- Dosages per plate



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## Plate Import / Export

Plate set setup information can be saved and reused using the import and export feature. Imports and exports are stored in clear text json format. This allows clients to export the data into their own processing systems or even edit the plates manually if the changes are very minor.

The export feature just requires a plate set to be selected and will save the file.

The Import feature receives a json export and asks the user to reassign some of the unique fields (plate set name and plate ids). It will populate the database based on the associated json.

## Updating Plate Set Status

Plate Sets can be updated from the main plate manager screen using the links next to the plate in the 'Action' column.

When a plate created, it reaches a preparing state. 'Mark prepared' will mark the plate as prepared and assign the marker as the preparer.

When the plate is prepared, the 'send to device' button will be available. Upon clicking it the state will change to 'in testing' and the marker will be identified as the tester.

When the plate is 'in testing' the upload analysis will become available. This link raises a popup allowing the user to upload the analysis provided by the machine. Upon upload, the state will change to 'tested'.

In the tested state, users will be able to reach the analysis interface using the 'view analysis' link.



## 6.2.2 Sample Screens / Story Board

### Plate Templates

The screenshot shows a web browser window with the address bar displaying 'http://www.we99.com'. The page title is 'A Web Page'. Below the browser window, there is a search bar with the text 'search'. The main content area is titled 'Plate Templates' and contains a table with the following data:

Name	Created by	Size	Select
Alan's Royal	alan orcherton	8x12 96 Well	<input type="checkbox"/>
Grandaddy Purple	Harry Humbold	8x12 96 Well	<input type="checkbox"/>
Freds Fantasy	Fead Willis	8x12 96 Well	<input type="checkbox"/>

Below the table, there is a sidebar with a '+ -' icon and a 'New Template' button. The sidebar also contains a list of options: '6 Well >', '24 Well >', '96 Well >', '384 Well >', 'Custom >', and 'Create From File >'.

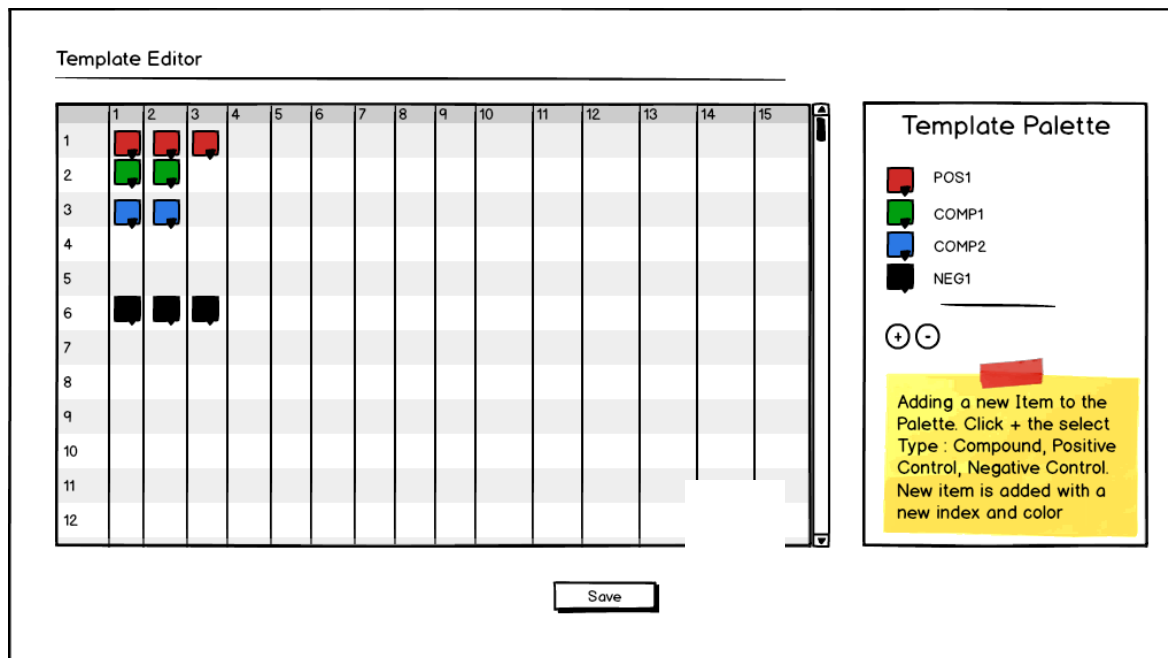
Plates are created from plate Templates.

- Scientists can create new templates using a template editor (next diagram).
- Scientists can also create new custom (rectangular) plates with the plate editor.
- A more common approach may be to upload existing CSV templates into the system.





## Plate Template Editor



The template editor allows scientist to create new plate templates. To each template the scientist adds various compound labels, and label for positive and negative controls.

When adding a new compounds, positive controls, or negative controls the scientist selects the “+” button in the Template palette. The scientist needs to select if s/he is adding a compound, negative, or positive control. Based on the selection a new item is added to the palette. Items get a new label and color following a convention e.g.

- Compound: COMP{number},
- Negative Control: NEG{number},
- Positive Control: POS{number}.

The Scientist can select the color from the palette and add the item to a particular well on the template.



A Web Page

http://www.we99.com

Q search

### Create Template From File - myExcelTemplate.csv

Name	Created by	Size	Status
John's_old_2003_template	John Wayne	96 Well	Unidentified Fields

#### Please Map Unidentified Fields

Label From File	Well Type
AB12	COMP1
SD52	COMP2
RT21	COMP3
myPositiveControl1	POS1
myNegativeControl1	NEG1

John Wayne's Convention  
Compound = COMP #  
Positive Control = POS #  
Negative Control = NEG #

Templates can be imported from file. The system will assign labels based on a default or user defined convention. If there are fields in the file that do not match the convention the user will be asked to map the label from the file to the convention. System needs to know if the label is a unique compound, a positive control, or a negative control

Save Save and View

Another option for creating template will be to create a template from a CSV file. Scientists may have a number of templates for previous experiments in excel format that can be exported to a CSV file.

We99 propose importing the CSV file to create templates using a labeling convention. Templates need to identify unique compounds, positive controls, and negative controls. A sample convention could be as follows:

- Compounds: COMP{Number}
- Positive Controls: POS{Number}
- Negative Controls: NEG{Number}

If the CSV file contains labels following the convention the import will be extremely simple. All the Scientist would have to do is provide a name for the imported template and save it to the database.

If the CSV file contains labels that do not follow the convention, then the Scientist will be prompted to map the unidentified fields to labels that follow the convention.

This concept could be extended to allow Scientists to have their own labeling convention for plate templates allowing them to import existing CSV templates easily.



WE99

http://www.we99isAwesome.com

### Tell us about your exciting experiment!

What is the name of your plate set:

What Type of plates are you using:

Need more plates? [Order more...](#)

Plate Type: Philosopher's Stone VX30. (Corning : Magic Rock)  
Rows: 20  
Cols: 10

What Template are you using:

**For this template you need:**

**3 Compounds**

Compounds
1. BlueJay45
2. TRXMarkIII

+ -

**1 Positive Controls**

Positive Controls
1. Proactiv

+ -

**1 Negative Controls**

Negative Controls
1. Brine

+ -

The scientist then creates plates sets for the experiment. During this stage the Scientist needs to map the real compounds and controls that are to be on the plate to compound and control labels on the template.

- The scientist provides a name for the plate set.
- Then selects the type of plate to be used (see ordering diagram). The plate type contains details regarding the actual physical plate to be used e.g. manufacturer, material etc.
- The scientist then selects the template to be used for the plate set.
- The system will determine number of compounds, positive controls and negative controls that need to be specified for the plate.
- The scientist adds the compounds and controls using a dropdown or picker from available compounds and controls stored in the database.
- Once complete the scientist selects continue



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Just a few more questions ...

How many Plates?

Initial Dose

Dilution Factor

Plate Set

Change the dosages for each plate if required

Plate	ID/Barcode	Dosage	UOM	Adjust Doses on Plate
1	XV12345	20	$\mu$ M	<input type="button" value="Adjust on Plate"/>
2				
3				











Option to adjust plate dose here, if possible UOM Should be both typeable and dropdown selection

The next step is to apply the dosage of the compounds for the plate. There are two ways to create dose response plates in the system.

1. Multiple plates each plate with diluted compounds
  - The scientist selects the number of plates.
  - Then selects the initial concentration.
  - Then specifies a dilution factor.
  - The system generates plates with different concentrations of compounds on each plate
  - The system will generate a unique ID for the plates, the scientist may override this to match any barcode ID that the physical plate may possess.
2. Single plate with multiple concentrations of compounds.
  - The scientist can choose an initial dose and no dilution factor
  - The scientist can then select "Adjust on Plate" to access the Plate Dose Editor







Plate Dose Editor: Plate ID XV12345

	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15
1															
2															
3															
4															
5															
6															
7															
8															
9															
10															
11															
12															

Type in Number on each compound to adjust dose for compounds on the same plate

Plate Key

-  Proactiv
-  BlueJay45
-  TRXMarkIII
-  Brine

Unit Of Measure

Save

- Using the plate dose editor the scientist can alter the concentration of each compound by changing the dosage value on each compound.

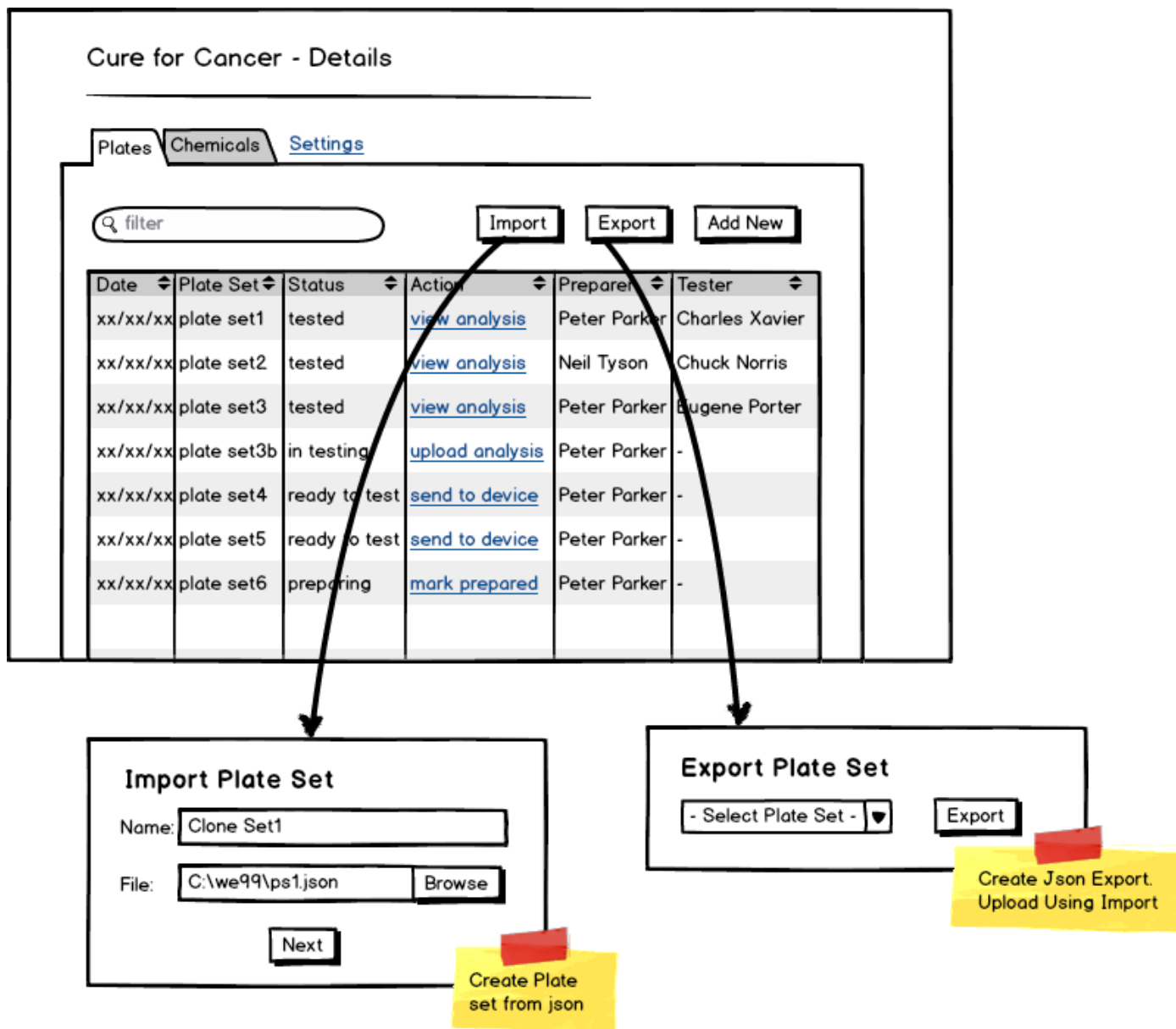


Plate Sets can be exported from the experiment manager into various formats.  
This will allow the scientist to create the physical plate to the specification created in the system.



### Plate Type Ordering Information

	Model	Manufacturer	Material	Columns	Rows	Dimensions
<a href="#">order</a>	X	Acme	Plastic	15	10	15cm x 10cm x 1cm
<a href="#">order</a>	Y	Acme	Glass	10	5	20cm x 10cm x 2cm
<a href="#">order</a>	Z	Acme	Plastic	150	100	300mm x 50mm x 5mm
<a href="#">order</a>	Philosopher's Stone VX3	Corning	Magic Rock	20	10	20cm x 10cm x 5cm

Brings up a new page with ordering information or goes to the 'order link' associated with the plate type

Close

Information regarding the Physical plate inventory on hand is also stored within the system. Scientists may order more plates through external links.

### 6.2.3 Open Issues / Questions Plate Management

#	DESCRIPTION	NOTES
1	Ability to create plates of any dimension. does not account for non-rectangular (e.g., round) plates or 3d plates.	Robert confirmed, this is NOT required
2	Exporting plate specs. Is there a standard format? Do we need to handle with different parsers tied to equipment type?	
3	We should consider having add plate set wizard to produce a json document server side. then the import wizard and the add new plate wizard would both use the same call to populate the database	
4	Concept of "Direction" for doses. Is this something we need to record for templates?	



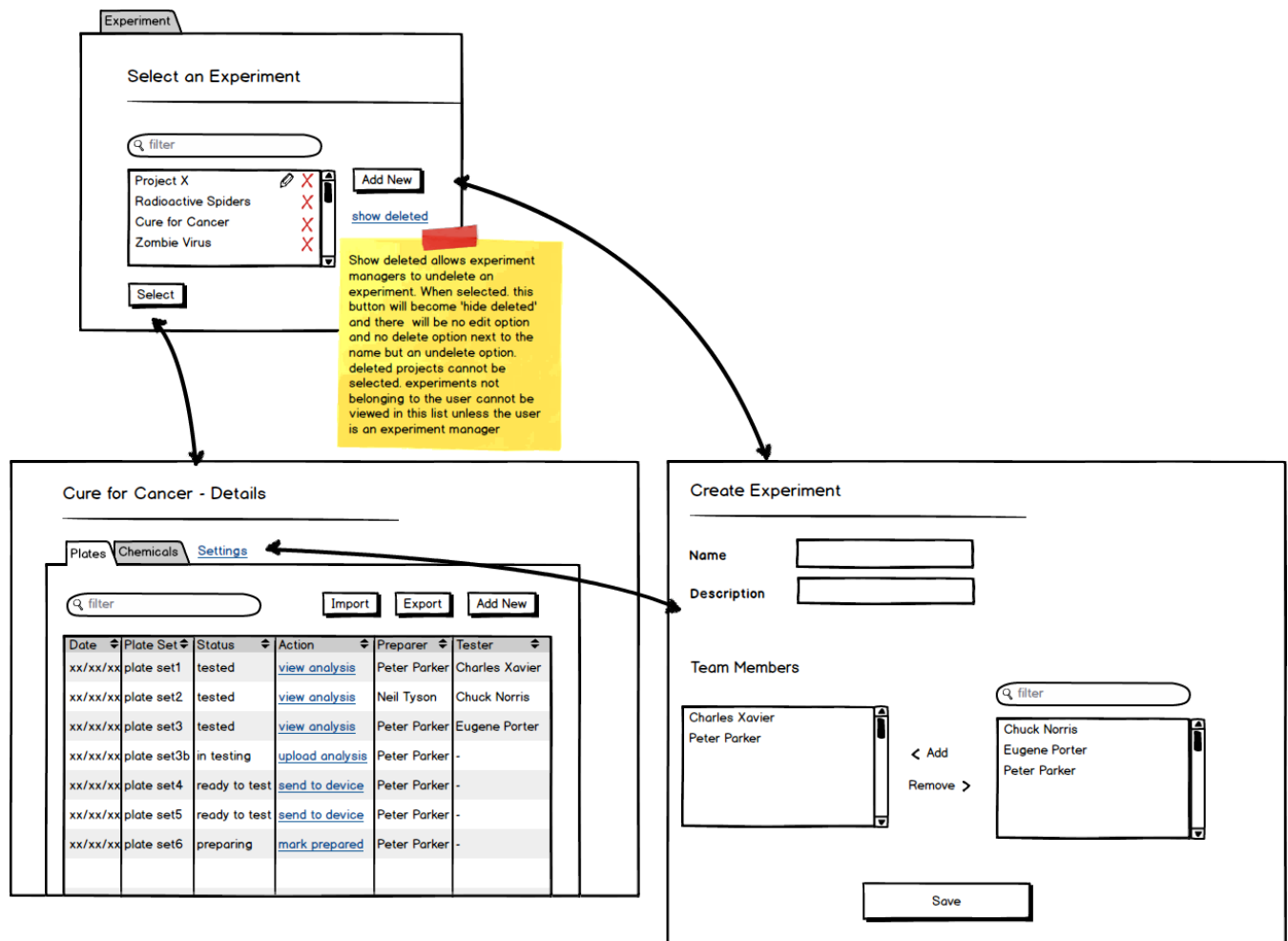
## 6.3 EXPERIMENT MANAGEMENT REQUIREMENTS

### 6.3.1 Overview

Experiments can be created and deleted by an administrative user. An experiment is identified with a unique name. A description of the experiment can also be included. An administrative user can also create teams of scientists for an experiment.

Experiments may use a single of a set of filled plates. The plates used in an experiment can be listed. A team member can add new plates to an experiment.

### 6.3.2 Sample Screens / Story Board







### 6.3.3 Open Issues / Questions Experiment Management

#	DESCRIPTION	NOTES
1	Adding Compounds. Is there a need to do this in a bulk import?	
2	note adding a team member removes them from the list of available team members	
3	<u>Might need Need a separate role and management view for 'mark as prepared'.</u>	<u>e.g., this is where a lab assistant could mark plates as prepared and the status is updated to the project.</u>



## 6.4 RESULTS ANALYSIS REQUIREMENTS

### 6.4.1 Overview

A plate will be analyzed by a machine - which will generate a structured text file as output. The user will upload the structured text file into the web application and select an appropriate parser to convert it into a result set. There is a large variety of output formats for analysis machines so the system must support the introduction of new text parsers. The results for the uploaded plates will be displayed to the user for validation in the form of a heat map.

While reviewing the results as a heat map the user will be able to read information about,

- 1) The results values.
- 2) The concentration of compounds in the wells.
- 3) The Z value of a particular well.
- 4) The compounds in the wells.
- 5) What type of well's are which (ex. EXP, POS, NEG).

Once an initial review of the wells is completed. The results can be saved to the system and the results as a whole can be QC'ed in a number of ways.

- 1) Individual plates can be brought up and reviewed for all of the above values.
- 2) The Z' value of each plate will be computed and displayed next to the plate.
- 3) The entire set of plates can be represented as a large heat map. A zoom in feature will allow the user to select specific areas of points to examine in greater detail.
- 4) Positive and Negative control wells, across all plates, over time can be represented to look at odd drifts in controls over time.

In either stage of the above process specific wells, or entire plates, can be tagged for exclusion due to unreliability.

Finally the results associated with the plate can be reviewed as scatter plots with a fitted dose response curves. The dosage amount associated with each well will be normalized by default. However the end user will be able to plug in normalization functions if they so wished. The dose response chart will be interactive and allow for the removal of specific points of data – and an adjustment of the fitted curve dynamically.

#### Algorithms –

A number of computations were mentioned above – and more may follow. So this section describes the algorithms that should be used for calculation.

**The Z value –** <http://en.wikipedia.org/wiki/Z-test>



$$z = \frac{\bar{x} - \Delta}{\frac{\sigma}{\sqrt{n}}}$$

This is calculated for each well on a plate. Where  $\bar{x}$  is the sample mean,  $\Delta$  is a specified value to be tested,  $\sigma$  is the population standard deviation, and  $n$  is the size of the sample.

**The Z' Factor** – <http://en.wikipedia.org/wiki/Z-factor>

$$\text{Z-factor} = 1 - \frac{3(\sigma_p + \sigma_n)}{|\mu_p - \mu_n|}.$$

This is calculated on the plate level. The Z-factor is defined in terms of four parameters: the means ( $\mu$ ) and standard deviations ( $\sigma$ ) of both the positive (p) and negative (n) controls ( $\mu_p$ ,  $\sigma_p$ , and  $\mu_n$ ,  $\sigma_n$ ).

**Curve Fitting** – [http://en.wikipedia.org/wiki/Curve\\_fitting](http://en.wikipedia.org/wiki/Curve_fitting)

This is calculated for a given compound across an entire experiment. This will use a polynomial fitting algorithm which fits the included points to a 3<sup>rd</sup> degree polynomial.

**Dosage Normalization** –

This is calculated on the dosage amount of a particular compound. The dosage amount is normalized using a logarithm function.

## 6.4.2 Sample Screens / Story Boards

### Story 1 – Uploading results to the system.

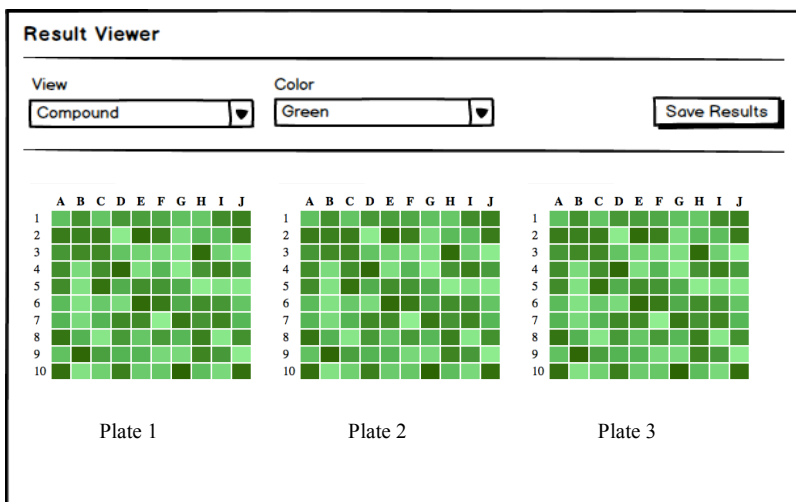
Step 1 – Select a file and an algorithm to upload.

Result File Uploader

File Name

Processing Algorithm Loader Algorithm 1 ▼

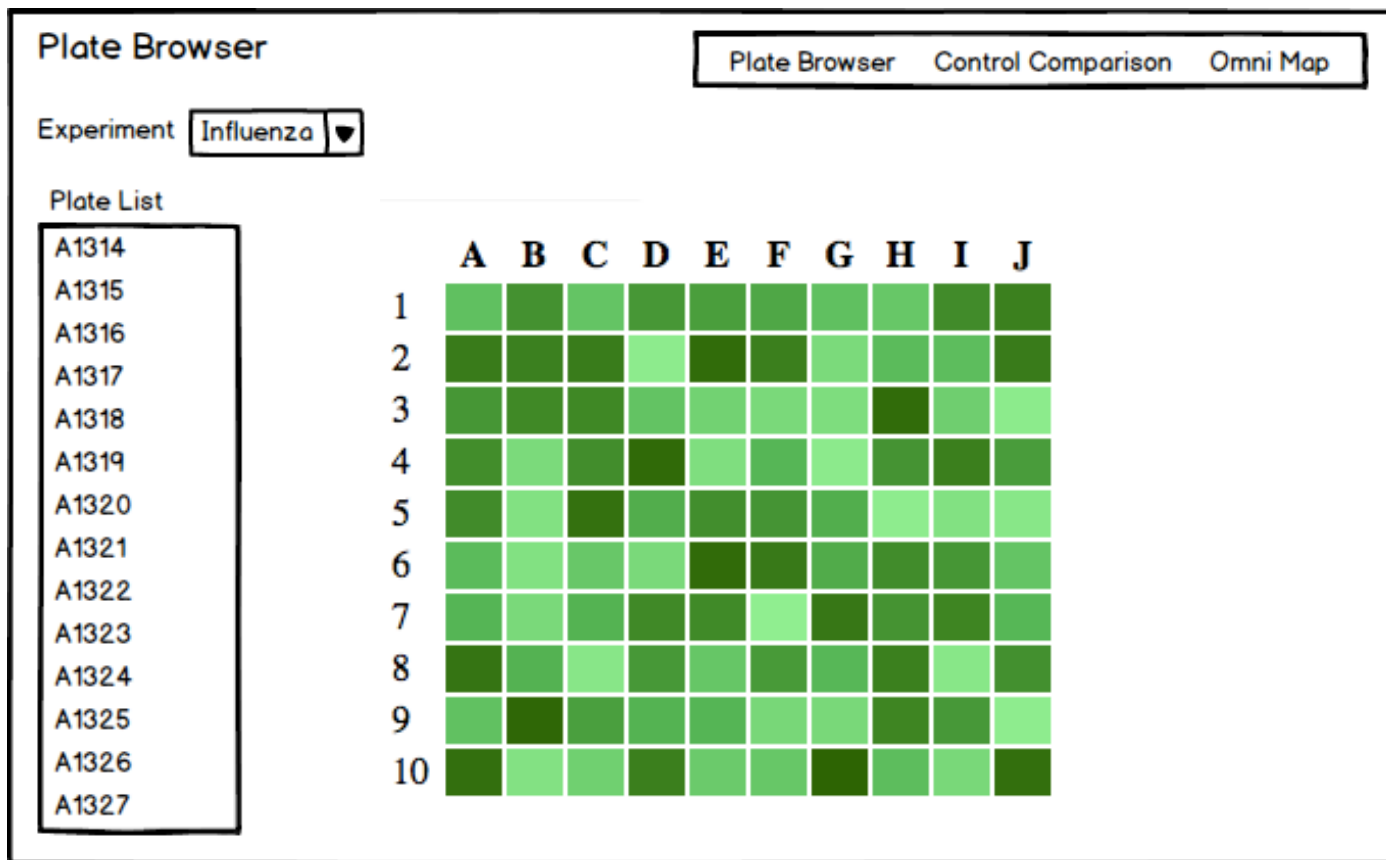
Step 2 – Review the results and save them if they look appropriate.



Step 2 – Initial review the results and save them if they look appropriate.

## Story 2 – Reviewing uploaded results for quality.

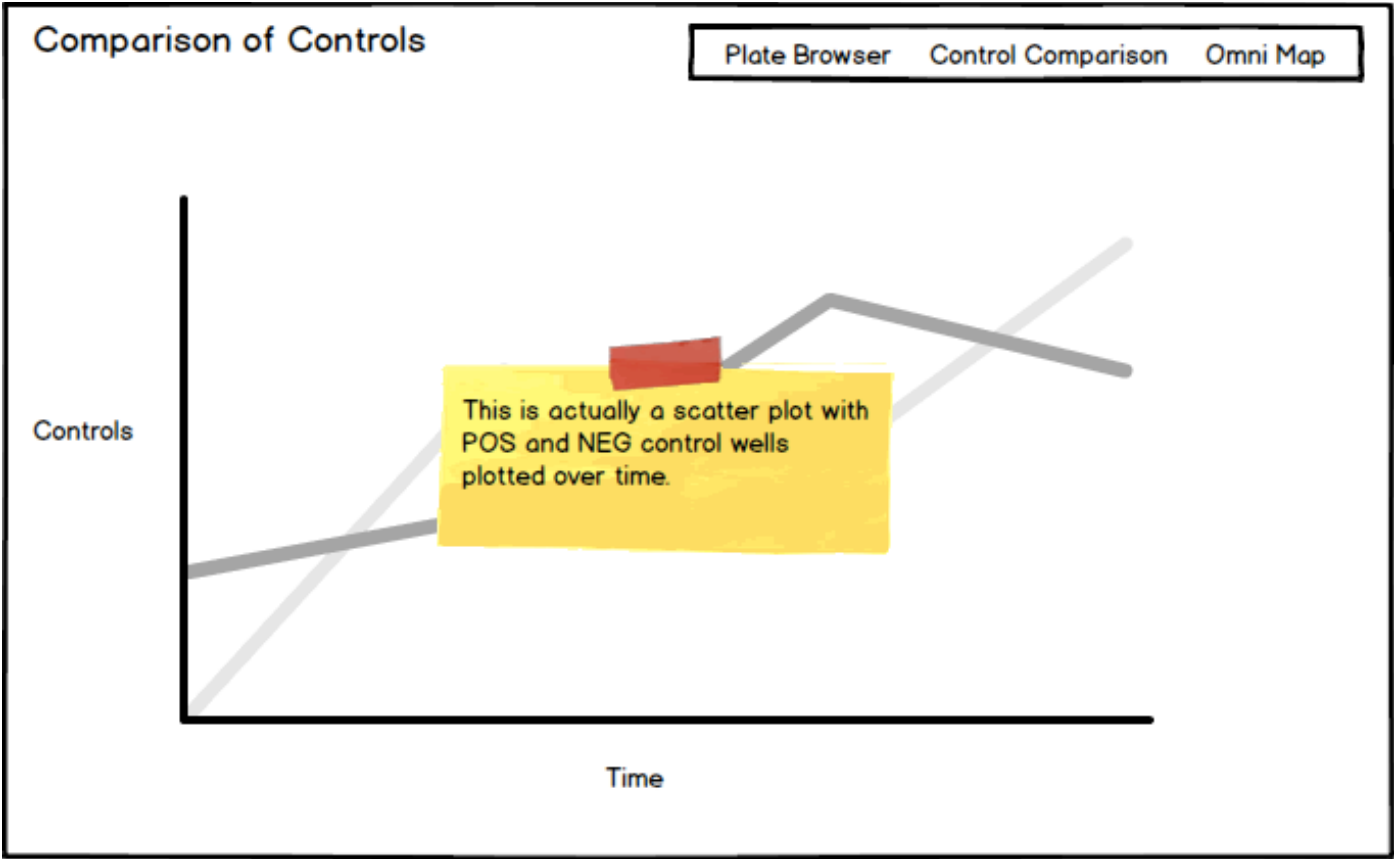
Step 1 – See the list of all plates in an experiment with associated plate level statistics.







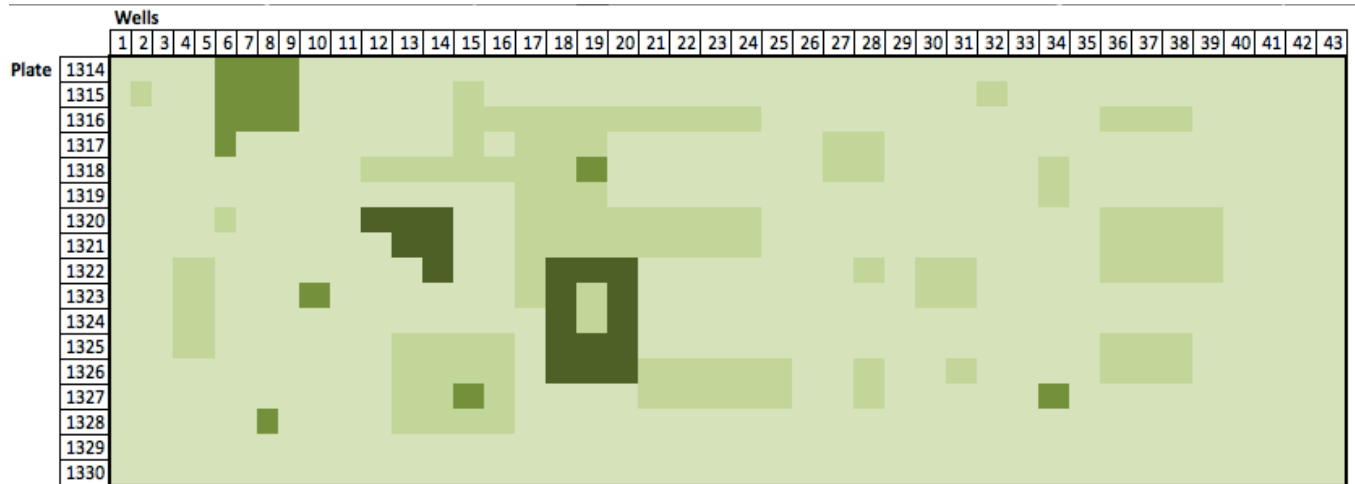
Step 2 – Review time series analysis of control wells.



Step 3 – Review the map of all wells on plates.



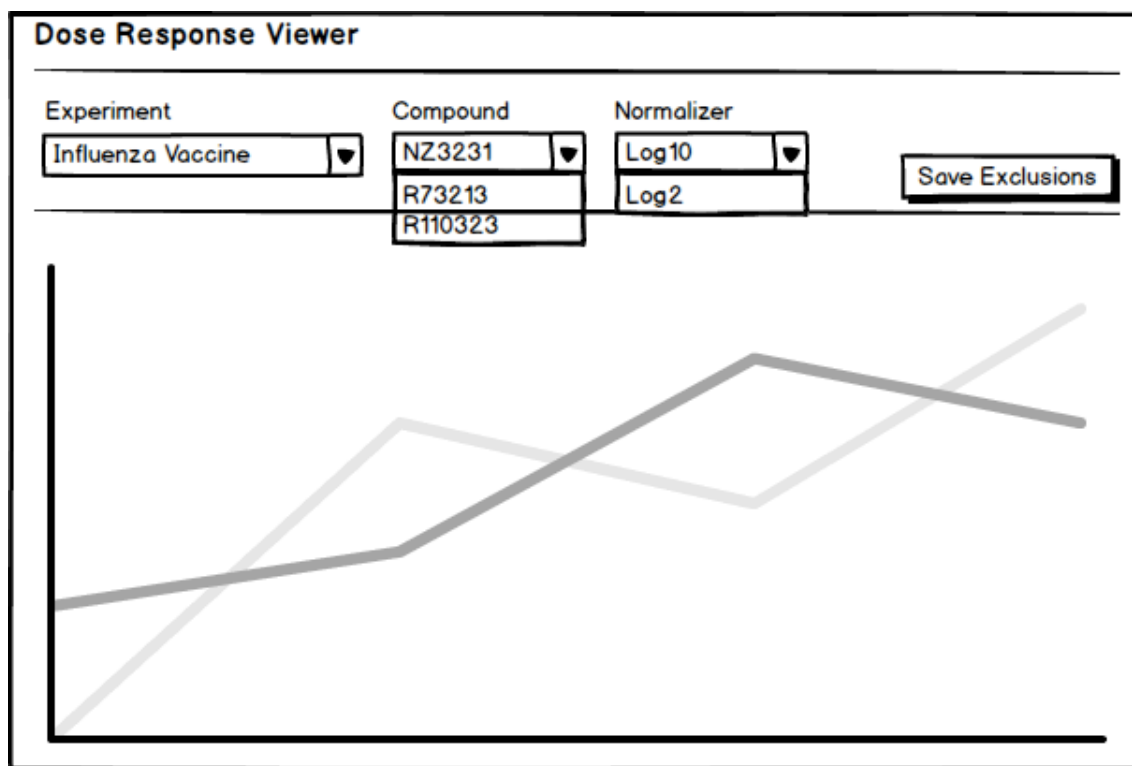
## Omni Map

[Plate Browser](#)[Control Comparison](#)[Omni Map](#)



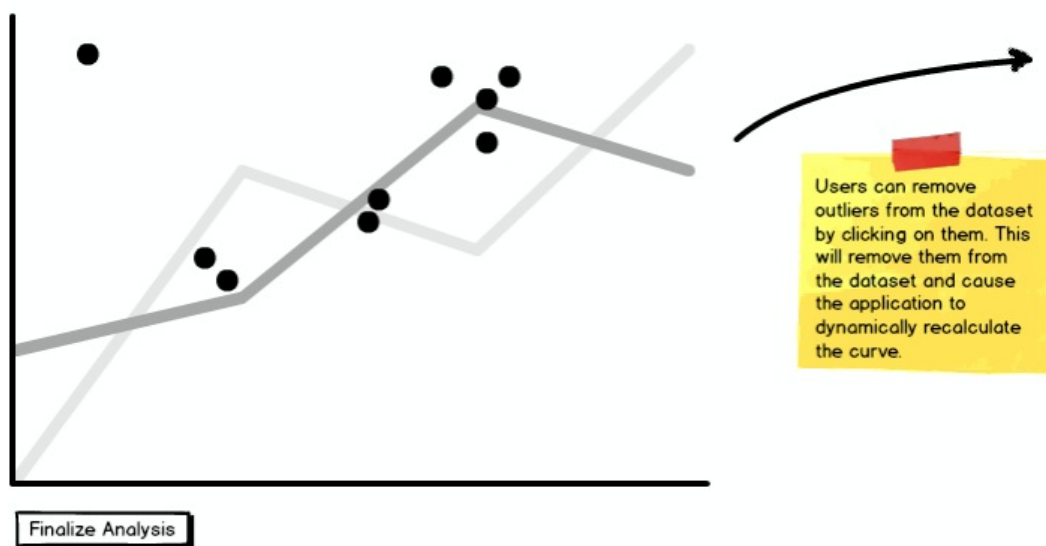
### Story 3 – Reviewing dose response curves.

Step 1 – Select an experiment, a compound, and a normalization function (if needed).



Step 2 – Review scatter plot and dose response curve for the compound then remove data points that are outliers to result in a cleaner result set.

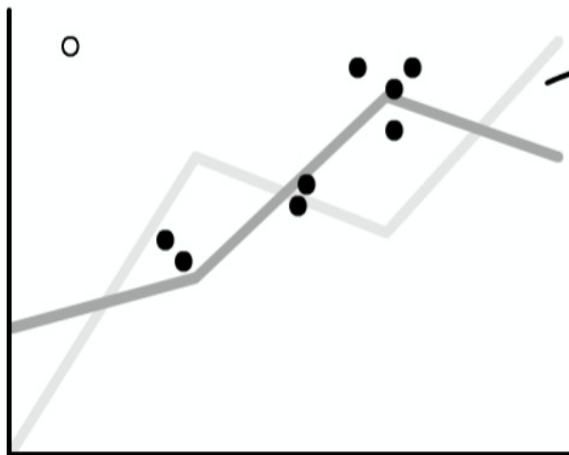
## Data Analysis







## Data Analysis



\* Modifications made to dataset

Finalize Analysis

When the user clicks "Finalize Analysis", they are brought to this screen. If outliers have been removed, they are listed in the table and the user is required to provide a justification in the text box. When user clicks save, the removed outliers list and the justification text is saved as the well as the analysis results in the database.

## Data Analysis

Analysis Name:

My Awesome Results

Removed Outliers:

Well #^	Concentration	Value
54	0.5	44
71	1.0	660
98	2.5	99

Justification for Outlier Removal (required):

Spilled coffee on right side of plate. Bad day

Save Analysis

### 6.4.3 Open Issues / Questions Results Analysis

#	DESCRIPTION	NOTES
1	When reviewing the result in the form of a heat map is there any workflow for processing the results beyond saving. For example -- if a plate looks like it has issues should the system allow them to flag the plate as problematic, etc?	



## 6.5 SPECIAL FEATURE PROPOSALS

### 6.5.1 Audit Trail

We could track whenever a Plate / Well / Dose / Experiment changes. This would allow users to know when something in the system has changed and may be necessary for internal reporting or perhaps regulatory agency requirements.

Assuming that we have some type of relational database, we should hook into the changes at the transaction level so the audit trail is captured in the same atomic operation as the data is changed. Note: there's a good library from Hibernate for this called Envers.

#### 6.5.1.1 Sample Screens / Story Boards

### Audit Logs

User	Event Type	Action	Experiment	Timestamp
George Washington	CREATE PLATE TEMPLATE	Created 'Acme Corp' Template	-	January 10 2015 4:00 AM
Abe Lincoln	CREATE PLATE	Created 'Abe's Compound X' plate	Geronimo	July 12 2014 6:00 PM
Thomas Jefferson	IMPORT RESULT	Imported file 'Output.dat'	Gamma	October 21 2014 9:00 AM
John Adams	SAVE ANALYSIS	Saved linear regression as 'My results'	Delta	Febuary 15 2015 7:00 AM
Abe Lincoln	DELETE PLATE TEMPLATE	Deleted template 'TEST123'	-	January 12 2014 6:00 AM

Export Logs

Export will allow the user to download audit logs as a csv file for backup or further analysis



## 6.5.2 Open Issues / Questions Special Features

#	DESCRIPTION	NOTES
1	For the data analysis portion of this they are represented as scatter plots. What should the x and the y axis of the scatter plot be?	
2	For the data analysis portion how should the result be filtered. Would people want to look at a specific compound across an entire experiment? Only look at a single plate at a time? Review all plates in an experiment?	

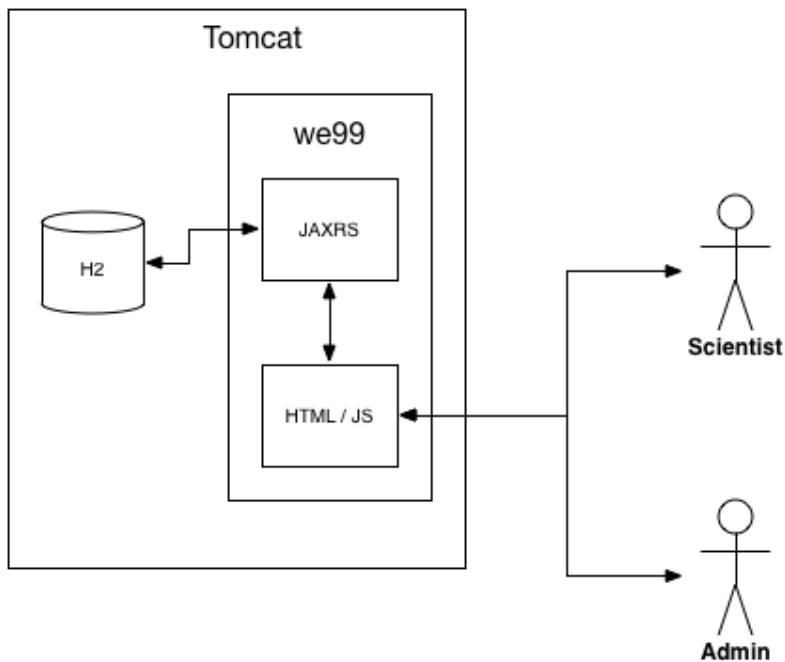


## 7. HIGH LEVEL DESIGN

### 7.1 ARCHITECTURE

#### 7.1.1 Introduction

Our overall vision is to create a web application that can assist scientists with managing the data associated with their experiments. We have decided to develop our project as a Java web application. A web application will easily allow multiple users to access our program from whatever internet-capable device they have available, such a workstation or a mobile device. Our team has significant experience in the Java development environment, so choosing a Java-based framework was considered the most efficient choice.



As a standard web app, we expect to see a Model, View, and Controller. This pattern is represented in the diagram where the model is the database that stores all of the data for the plates and the associated experiments. The controller part is the REST web services that provide all of the operations necessary to create and modify experiment data. Finally, the presentation layer is implemented in a HTML and Javascript and interacts with the controllers through standard AJAX/HTTP calls.



### 7.1.2 Model

The data components in the application are modeled as plain Java Beans. The beans are named using domain in accordance with domain specific terms in order to better associate the data models with real world tasks needed to be done by the domain experts.

The domain model uses the standard JPA annotations in order to be able to create a declarative persistence layer for the application. While this persistence layer is primarily focused on a relational database implementation, it's not strictly required. It's conceivable for the customer to swap the persistence layer out at a future point in favor of another implementation without having to undertake a complete rewrite. This is the declarative nature of the persistence model. At best, it is loosely coupled to a relational database.

### 7.1.3 Controller

The controller portion of the application is implemented via REST services. The service classes are implemented using the JAXRS standard specification. As such, they will run in any JAXRS compliant container. Furthermore, the services can be exercised by any REST client capable of producing the documented JSON and sending it to the documented URI's.

One added benefit of using the standard JAXRS annotations is that we can leverage the capabilities provided by one popular implementation (Apache CXF) in order to provide strongly typed client API's in order to assist with future integrations. Obviously future integrators are free to choose any library that suits their needs, but if they happen to choose Apache CXF or a similarly compatible library, then they can use our interfaces directly in order to generate dynamic proxies to invoke our services.

### 7.1.4 View

The view component of the application is an application based on Angular JS. This framework is an open source HTML 5 framework from Google. The application will have knowledge of the set of permissions in the application and render the UI accordingly. It will exercise the service components and allow the user to interact with the plates and experiment results. All of the interactions from the client to the server will be accomplished through the standard REST service interfaces.

### 7.1.5 Components

Both the front-end and back-end are design by contract. The REST services provide sufficient annotations and documentations to enable the implementations of a full application. The front-end will extract each of its service calls to a well defined service interface that could be swapped for a different implementation.



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### 7.1.6 Database

The default database vendor is the H2 project. This is a SQL 92 compliant database that has wide support across major libraries like Hibernate and the Java Persistence Architecture. The key thing to consider with a database vendor is whether you're locked into the database vendor through some vendor specific API. Specifically, is there some touch point with the vendor where the host application is interacting with the DB host in an interface that is not part of a standard interface like JDBC or JPA.

Being vendor agnostic is a key design principle with WE99. We'll avoid any vendor specific queries or other enhancements. No vendor specific SQL, no stored procedures, and no assumptions about the platform executing our queries. This vendor agnostic layer is provided largely for free via the Java Persistence Architecture API. We can encode our DB access into our entities via annotations and an abstract query language that has late binding to a vendor specific query language at running.

The initial deployment of WE99 is designed to scale vertically as opposed to horizontally. This could change by rethinking the persistence layer to introduce an abstraction layer for JPA. This indirection would make it easier to swap out a traditional RDBMS with a NoSQL approach if needed. While there's still some work to do to support this, it's isolated to the service implementations and not part of the core interface with the presentation layer.



## 8. PROJECT PLANNING AND ESTIMATING

### 8.1 ITERATIONS

#### 8.1.1 Overview

We99 has planned the project based on the Agile Methodology. The project is broken down into iterations. The We99 team has planned 10 iterations, each iteration being 1 week in duration. The iteration starts on the Thursday of each week and closes on the following Wednesday. Every iteration the project team will be delivering several units of functionality defined by stories.

#### 8.1.2 Iteration Calendar

The table below specifies the iteration calendar for the We99 Team

Iteration Number	Open Date	Close Date	Notes
1	March 5 <sup>th</sup> 2015	March 11 <sup>th</sup> 2015	Milestone1 Presentation March 5th
2	March 12 <sup>th</sup> 2015	March 18 <sup>th</sup> 2015	
3	March 19 <sup>th</sup> 2015	March 25 <sup>th</sup> 2015	Spring Break March 15 <sup>th</sup> -22nd
4	March 26 <sup>th</sup> 2015	April 1st 2015	
5	April 2 <sup>nd</sup> 2015	April 8 <sup>th</sup> 2015	Milestone 2 Presentation April 9th
6	April 9 <sup>th</sup> 2015	April 15 <sup>th</sup> 2015	
7	April 16 <sup>th</sup> 2015	April 22 <sup>nd</sup> 2015	
8	April 23 <sup>rd</sup> 2015	April 29 <sup>th</sup> 2015	
9	April 30 <sup>th</sup> 2015	May 6 <sup>th</sup> 2015	Milestone 3 Presentation May 7th
10	May 7 <sup>th</sup> 2015	May 13 <sup>th</sup> 2015	Final Report May 9 Presentation Faculty Committee May 14



## 8.2 STORIES AND ESTIMATES

The We99 are using Atlassian JIRA to manage and track project stories.  
<https://westeast99.atlassian.net/secure/Dashboard.jspa>.

As of march 2<sup>nd</sup>, The team have 30 stories in the backlog. Each story is sized with an attribute called story points. Each point represents a team man hours. The initial rough estimates predict that the team will require 360 points (man hours) to complete the project. Our initial estimates are rough, so doubling the estimate out estimated (man hours) for the project is 720 Hours. The team has a capacity of 750 hours for the project.

The We99 team predicts that they should be able to complete the scope of work for the project within the allotted timeframe.

At the close of every iteration the team will re-compute its “velocity”. That is the number of story points it is averaging per iteration. This is an iterative process and the team will adjust its estimates accordingly.

## 8.3 MAIN DELIVERABLES

Milestone 2 – April 9<sup>th</sup>. Close of iteration 5

- Predicted Functionality completed
- Domain Model Completed
- Web Service Interfaces specified
- Web Services Implemented for Plate Editor
- Web Services Implemented for Plate Quality Control
- Web Services Implemented for
- Plate Template Editor for the creating of Dose Response Plates
- Front End Quality Control of Plates

Milestone 3 – May 7<sup>th</sup>. Close of iteration 9





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## 9. PROJECT RISKS

### 9.1 RISKS IDENTIFIED



## 10. APPENDIX STORY BACKLOG MARCH 2ND

Issue Type	Key	Summary
Story	<a href="#">WE-17</a>	Update Report and Presentation with Architecture and Web Service Info
Story	<a href="#">WE-4</a>	Import CSV Plate Maps
Story	<a href="#">WE-19</a>	Update Report and Presentation with Front End Stack
Story	<a href="#">WE-18</a>	Update Report and Presentation with Results Analysis Requirements
Story	<a href="#">WE-20</a>	Update Report and Presentation with details of the Plate Map Editor
Story	<a href="#">WE-3</a>	Basic LogIn Logout
Story	<a href="#">WE-6</a>	Implement Standard Equations for Dose Response Curves
Story	<a href="#">WE-7</a>	Front End - Support User-Specified fit Equations for Dose Response curves
Story	<a href="#">WE-8</a>	BackEnd Services to support HeatMaps for plate results
Story	<a href="#">WE-11</a>	Back End Services to Compute Z and Z' for QC'ing Plates
Story	<a href="#">WE-12</a>	Plate Map Editor
Story	<a href="#">WE-13</a>	Import and parse test results
Story	<a href="#">WE-14</a>	Design Domain Model
Story	<a href="#">WE-15</a>	Define Web Service Interfaces
Story	<a href="#">WE-24</a>	BackEnd Results Parser - to parse and store result files
Story	<a href="#">WE-25</a>	Front End Results File picker - select results file to parse
Story	<a href="#">WE-26</a>	Front End - Display Single Heat Map for a Single plate
Story	<a href="#">WE-27</a>	Front End - Display Multiple Heat Maps for Experiment
Story	<a href="#">WE-28</a>	Front End - Display Single Dose Response Curve
Story	<a href="#">WE-29</a>	Front End - Knock out Values on Dose Response Curve and Refit
Story	<a href="#">WE-30</a>	Front End - Display plate statistics, and control well statistics across all the plates for all teams



Story	<a href="#">WE-31</a>	Back End - Support Services for plate statistics, and control well statistics across all the plates for all teams
Story	<a href="#">WE-32</a>	Front End - Support Combination views of Dose response curves
Story	<a href="#">WE-33</a>	Front End - Specify values for fit parameters for all or a Selection of Dose response curves
Story	<a href="#">WE-34</a>	Front End - Select %effect ranges for to plot non, partial and full responders Dose response curves
Story	<a href="#">WE-35</a>	Back End - Store Raw Results file for reproduction of analysis
Story	<a href="#">WE-36</a>	Back End - Define negative and positive controls so all normalized results are in terms of a percent effect
Story	<a href="#">WE-37</a>	Back End - Store control data
Story	<a href="#">WE-38</a>	Back End - Store SVG plots transforms and meta data for plots
Story	<a href="#">WE-39</a>	Back End - Audit and store all Knock out values during dose response analysis