

APPROVED

By Jean Gildner at 11:46 am, Jul 19, 2016

From: [Amin, Pankaj \(Pete\)](#)
To: [Gildner, Jean](#)
Cc: [Lorenzo, Anthony](#); [Bobo, Qiao](#); [Thomas, John](#)
Subject: RE: Information request questions
Date: Friday, May 27, 2016 12:02:12 PM

I have following Information request (IR questions) for Vericel BLA 125603 regarding Facility section 3.2.A):

Question #1

Section 1.7 describe the prevention of contamination and cross contamination. Please provide following additional information:

- Please provide a description of the sanitizer effectiveness assessment performed for (b) (4) used in (b) (4) areas.
- Please describe room clearance and changeover procedure in place to prevent cross contamination.
- Please provide a narrative summary of how the manufacturing operations and storage of other licensed products are segregated from MACI production and storage areas.
- Please describe equipment cleaning/sanitization procedures used in (b) (4) for product contact equipment.

Question #2

Section 1.1.6 describes Environmental monitoring program. Please provide following additional information for MACI production areas:

- Please provide a description of your routine monitoring program including the testing performed, frequency of testing, alert and action limits and sampling locations.
- Please provide the environmental monitoring qualification data for the new cell therapy suite (b) (4) including BSC hoods.
- Please provide summary of the most recent three months routine environmental monitoring data.

Question #3

Section 1.3.1(product contact equipment) states that (b) (4) validation and routine (b) (4) verification audits are performed to confirm the custom made membrane loading units are suitable for use. The materials are sourced from approved vendors and (b) (4) or (b) (4). Please provide following information for (b) (4), membrane loading unit, forceps, scalpel and primary container and (b) (4) :

- Name of approved vendors and summary of vendor qualification completed
- How (b) (4) data provided in the certificate of analysis were validated
- Describe acceptance criteria for the product contact equipment
- Describe your (b) (4) verification audit and current status of these audit

Question #4

Section 1.2.3 (b) (4) and section 1.2.4 (b) (4) describe (b) (4).
Please provide following additional information:

- Describe approved vendor name and vendor qualification summary (brief)
- Copy of certificate of analysis (vendor provided)
- Please explain why you did not establish acceptance criteria for (b) (4) for the (b) (4) system and clarify if you routinely monitor the (b) (4).

Question #5

Section 1.2.1 describe Heating Ventilation and Air Conditioning (HVAC). Please provide following additional information:

- Please provide a diagram that shows the location of air handling units and rooms they service, locations of the terminal HEPA filters, and an enlargement of the area. classification diagram for the (b) (4) production areas.
- Please describe total number of air handling units that serve (b) (4) including clarification if air is recirculated or once through.
- Please provide an enlargement of the area pressurization diagram that shows the production areas on (b) (4) floor.
- Please provide HVAC qualification protocol and report (OQ/PQ) for the MACI production areas.
- Please provide the clean room (cell therapy suite- (b) (4)) qualification summary.
- Please provide Biological safety hood (BSC) qualification summary for the (b) (4)

Question #6

Section 1.2.2 describes (b) (4) purified water system. Please provide following additional information:

- Briefly summarize the purified water system description and qualification/requalification
- Summary of validated acceptance criteria
- Routine monitoring frequencies, acceptance criteria

Thanks

Pete Amin

From: Gildner, Jean
Sent: Thursday, May 26, 2016 1:19 PM
To: Amin, Pankaj (Pete)
Subject: RE: Information request questions

Hi Pete,

I will be looking for your questions tomorrow.

Thanks,

Jean

From: Amin, Pankaj (Pete)
Sent: Wednesday, May 25, 2016 4:03 PM
To: Thomas, John; Gildner, Jean
Subject: Information request questions

As part of my facility review of Vericel BLA, I have list of IR questions that I will send you by Friday. Please note that all my IR questions are relate to getting more clarification/ information and none of these questions are show stopper for this BLA (will not be part of next week briefing meeting).

Thanks
Pete