GMP Site Quality Self-Assessment

based on

Rx-360 Supplier Assessment Questionnaire

Module 2, Site Specific Information

Relevant for

Sigma-Aldrich, Inc.
3300 South Second Street (Cherokee)
St. Louis, MO 63118, USA
An affiliate of Merck KGaA, Darmstadt, Germany

The site self-assessment covers our quality management system for the following regulated applications:
- Manufacturing of pharma raw materials

The site also processes products which do not fall under any of the above mentioned regulated areas. For these products, other quality standards apply. For details, please refer to our Non-GMP Quality Site Self-Assessment.

Rx 360 An International Pharmaceutical Supply Chain Consortium

Merck KGaA, Darmstadt, Germany is an active member of the Rx 360 Consortium.

As a trusted partner of our customers, we deliver quality - always.

Merck KGaA
Corporation with General Partners
Frankfurter Str. 250
64293 Darmstadt, Germany

Information

This document is based on the Rx-360 Consortium's Supplier Assessment Questionnaire template, Module 2.
The contents of this questionnaire are built on the Rx-360 questionnaire version 2.0 intact with no question added or deleted.
Rx-360's CEO/COO gave permission to Life Science to use the Rx-360 logo.

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Rx-360 Supplier Assessment Questionnaire : Site-Specific Information

☐ Please check here if additional documents are attached.

SECTION 1. General Site Information	
1.1	Site or Facility-Specific Name: MilliporeSigma
1.2	Address: 3300 S. Second St., St. Louis, MO 63118 USA GPS Coordinates: GPS Coordinates: guest entry at 3360 S. Second St. is Lat: 38° 35' 27.9852", Long: -90° 12' 48.204"
1.3	Phone: 314-286-6600
1.4	Email: Please refer to your responsible Sales representative
1.5	Fax: N/A
1.6	Website: milliporesigma.com

SECTION 2. General Site Operating Information	
2.1	What year did the site start operating? Site in 1986 with upgrade to GMP in 1995.
2.2	What is the primary activity of the site? (e.g. manufacturing, distribution, etc.) Manufacturing, process and analytical development, testing and packaging
2.3	To which, if any, subdivision of the parent company does the site belong? Merck KGaA, Darmstadt, Germany

SECTION 2. General Site Operating Information		
2.4	Size of site (in sq. ft. or m.): 400,000 sqft Site	
2.5	Please list or attach the normal hours/schedule of the facilities, including shutdown dates (if applicable): Bio-Organics operates 24 hours a day, 7 days a week, 365 days a year. Bioconjugation and Protein Purification facilities operate 24 hours a day 5 days a week when in production, otherwise M-F first shift. Maintenance shut downs occur within individual departments twice a year, with staggered timeframes for each department	
2.6	Total number of employees on site: 380	
2.7	Total number of employees in Quality: 78	
2.8	Total number of employees in Manufacturing: 57	
2.9	What quality management system is utilized on site? ☒ ISO 9001 ☐ ISO 13485 ☒ 21 CFR Part 210/211 ☐ 21 CFR Part 820 ☐ European GMP, Eudralex Volume 4 Part I ☐ European GMP, Eudralex Volume 4 Part II ☒ ICH Q7 ☐ HACCP ☐ ISO 22000 ☐ Other Please describe: cGMP facilities and processes at Cherokee will not be associated with any ISO certification except of Maintenance Which Regulatory Initiatives does the site follow/comply with? ☒ REACH ☒ RoHs ☒ Ca Prop. 65 ☐ WEEE	
2.10	Does the company/site have an export license?	☒ Yes ☐ No ☐ N/A

SECTION 2. General Site Operating Information		
2.11	Is the site registered with any government regulatory agency (FDA registration, GMP certification, etc.)? ☒ Yes ☐ No ☐ N/A If yes, please specify. FDA registration 1937990 and USDA MO-TEC-0015	
2.12	By whom is the site inspected (regulatory or third party) and list inspections within the last three years: FDA, USDA, MFDS	
2.13	How often, as an annual average, is the site audited by customers or third parties? 20 times per year	
2.14	Has an Rx-360 audit been performed at this site? ☒ Yes ☐ No Please also state the date of the audit if applicable. Please refer to list on the internet for purchasing audit reports. Last RX-360 audit performed at the site in September 2021. http://rx-360.org/audit-programs/	
2.15	Are you willing to have Rx-360 conduct an audit on behalf of your customers according to the Rx-360 audit programs on your site? ☒ Yes ☐ No	
2.16	Are you willing to have your customers conduct audits on your site? ☒ Yes ☐ No	
2.17	Please list regulatory sanctions impacting the site within the last five years (i.e. warning letters, CEP suspension, import alerts, etc.): No sanctions to reference	
2.18	Does the site outsource any quality-related activity? ☒ Yes ☐ No ☐ N/A If answering yes, please specify the activities: Some testing is sourced to outside laboratories. Some calibrations activities and pest management.	
2.19	Please check the supplier controls in place for this facility:	
2.19a	Quality Agreements with Suppliers	☒ Yes ☐ No ☐ N/A
2.19b	Subcontractor Qualification/Audit Program	☒ Yes ☐ No ☐ N/A

SECTION 2. General Site Operating Information				
2.19c	Periodic Review of Supplier Performance	☒ Yes	☐ No	☐ N/A
2.19d	Supplier Feedback Program	☒ Yes	☐ No	☐ N/A
2.19e	Approved Material Supplier List	☒ Yes	☐ No	☐ N/A
2.19f	Approved Service Supplier List	☒ Yes	☐ No	☐ N/A
Additional comments: N/A				

SECTION 3. Objectionable Materials on Site				
3.1	Does the site or production plant produce, process or store any of the following:	**Yes**	**No**	**Not Applicable**
3.1a	Beta-Lactam Antibiotics	☐	☒	☐
3.1b	Steroids and/or hormones	☐	☒	☐
3.1c	High potency compounds	☒	☐	☐
3.1d	Materials of animal origin/Biologics	☒	☐	☐
3.1e	Live virus or micro-organism	☐	☒	☐
3.1f	Allergens	☐	☒	☐
3.1g	Genetically Modified Organisms (GMO)	☐	☒	☐
3.1h	Agrochemicals (Pesticides, Herbicides, Fungicides, etc.)	☐	☒	☐
3.1i	Other (Please specify): Section 3.1d - Cherokee no longer manufactures with animal origin components, however finished product animal containing components are stored in dedicated temperature units in an animal component containing facility.			

SECTION 4. Cross Contamination Control				
4.1	Are any of the following cross-contamination controls in place?	**Yes**	**No**	**Not Applicable**
4.1a	Dedicated Facilities	☒	☐	☐
4.1b	Access Controls	☒	☐	☐
4.1c	Dedicated Personnel	☒	☐	☐
4.1d	Dedicated Gowning	☒	☐	☐
4.1e	Procedural Controls	☒	☐	☐
4.1f	Other (please specify): N/A			

Additional Comments: N/A

	SECTION 5. Site Operating Policies			
		Yes	No	Not Applicable
5.1	Does the site utilize the following written policies, programs, or procedures?	☒	☐	☐
Site Specific:				
5.1a	Environmental, Health, and Safety	☒	☐	☐
5.1b	Facility Environmental Control Policy	☒	☐	☐
5.1c	General Facility Cleaning Procedures	☒	☐	☐
5.1c	Hygiene and Sterilization Procedures	☒	☐	☐
5.1d	Validated Equipment Cleaning Procedures	☒	☐	☐
5.1e	Preventative Maintenance Program/Procedures	☒	☐	☐
5.1f	Pest Control Program	☒	☐	☐
5.1g	Master Production Procedure	☒	☐	☐
Quality:				
5.1h	Quality Control/Quality Management Policy	☒	☐	☐
5.1i	Quality Manual	☒	☐	☐
5.1j	Periodic Product Quality Review	☒	☐	☐
5.1k	Master Validation Plan	☒	☐	☐
5.1l	Risk Assessment Program	☒	☐	☐
5.1m	Supplier Approval Procedure	☒	☐	☐
5.1n	Monitoring and Review of Approved Suppliers	☒	☐	☐
5.1o	Mechanism to Reduce Testing	☒	☐	☐
5.1p	Receiving Incoming Inspection	☒	☐	☐
5.1q	Change Control Procedures	☒	☐	☐
5.1r	Document Management Policy	☒	☐	☐
5.1s	Document Retention Policy	☒	☐	☐
5.1t	Change Notification Procedures for Clients	☒	☐	☐
5.1u	Control of Nonconforming Material	☒	☐	☐
5.1v	Deviation/Investigation Procedure	☒	☐	☐
5.1w	Out of Specification Policy and Procedure	☒	☐	☐
5.1x	Sampling Procedure/Sampling Plan	☒	☐	☐
5.1y	Raw Material Retention Program	☒	☐	☐
5.1z	CAPA Procedure	☒	☐	☐
5.1aa	Label Control and Accountability	☒	☐	☐
5.1bb	Product Release Procedure	☒	☐	☐
5.1cc	Employee Training Program	☒	☐	☐
5.1dd	Stability, Expiration, and Shelf-Life Program	☒	☐	☐
5.1ee	Product Retention Program	☒	☐	☐

5.1ff	Recall Procedure	☒	☐	☐
5.1gg	Customer Complaint Handling	☒	☐	☐
5.1hh	Equipment validation/qualification procedure	☒	☐	☐
5.1ii	Internal audit/self-inspection program procedure	☒	☐	☐
5.1jj	Site Security/Site Access Control Policies	☒	☐	☐
5.1kk	New Hire Program/Induction Program	☒	☐	☐
Business Continuity/Contingency Plan:				
5.1ll	Disaster Recovery Plan	☒	☐	☐
5.1mm	Pandemic Preparedness Plan	☒	☐	☐
5.1nn	Supply Chain Emergency Preparedness Plan	☒	☐	☐
5.1oo	Business Continuity/Contingency Plan	☒	☐	☐
5.1pp	Can the company provide a plan upon request? OR provide a short description below: Yes			

SECTION 6. Quality Assurance and Production				
		Yes	**No**	**Not Applicable**
6.1	Does the site have an independent and defined Quality Assurance/Quality Management Division?	☒	☐	☐
6.2	Does QA/QM have authority over the following:	☒	☐	☐
6.2a	Policies and procedures?	☒	☐	☐
6.2b	Review of documentation for release?	☒	☐	☐
6.2c	Release or rejection of incoming materials?	☒	☐	☐
6.3	Does QA/QM investigate and resolve quality complaints?	☒	☐	☐
6.4	Does QA/QM investigate and resolve internal deviations?	☒	☐	☐
6.5	Does the QA/QM have the authority to assign a disposition to materials?	☒	☐	☐
6.6	Does the QA/QM review manufacturing and testing records prior to release?	☒	☐	☐
6.7	Does the facility utilize computerized systems for managing GxP activities or data?	☒	☐	☐
6.8	Are relevant computerized systems 21 CFR part 11 and EU GMP annex 11 compliant?	☒	☐	☐
6.9	Does the site use statistical methods for consistency and uniformity?	☒	☐	☐
6.10	Does the site use controlled documents for following and recording manufacturing instructions?	☒	☐	☐
6.11	Does the company qualify and/or validate manufacturing procedures?	☒	☐	☐

SECTION 6. Quality Assurance and Production				
		Yes	**No**	**Not Applicable**
6.12	Is any environmental monitoring conducted in production/finishing areas?	☒	☐	☐
6.13	Does the site supply BSE/TSE declarations?	☒	☐	☐
6.14	Does the site supply a declaration of Elemental Impurities?	☒	☐	☐
6.15	Are ICH Q3C(R4) solvents used in the manufacturing process of supplied materials?	☒	☐	☐
6.16	Are stability studies carried out according to ICH guidance?	☒	☐	☐
6.17	Are solvents and mother liquor reused/recycled?	☐	☒	☐
6.18	Does the site have a process water treatment system?	☒	☐	☐
6.18a	Please check all that apply to the system: ☐ City/potable water ☐ Distilled water ☐ Dionized water ☒ Water for injection (WFI) ☒ Reverse Osmosis ☒ Clean steam ☒ Ultra-filtrated water (purified water) ☐ Other:			
6.19	Does the plant have a batch/lot system?	☒	☐	☐
6.19a	Is the system traceable?	☒	☐	☐
6.19b	Is it unique?	☒	☐	☐
6.19c	Is batch/lot manufacturing continuous?	☐	☒	☐
6.19d	Is manufacturing batch by batch?	☒	☐	☐
6.20	Does the site perform on-plant audits prior to approving critical GxP suppliers?	☒	☐	☐
6.21	Does the site audit critical GxP suppliers after initial approval?	☒	☐	☐
6.22	Does the site inspect incoming materials?	☒	☐	☐
6.23	Does the site test incoming materials to defined specifications?	☒	☐	☐
6.24	Does the site establish purchase specifications for raw materials?	☒	☐	☐
6.25	Is the equipment multi-use?	☒	☐	☐
6.26	Does the site qualify equipment installation?	☒	☐	☐
6.27	Does the site qualify equipment operation?	☒	☐	☐
6.28	Does the site qualify equipment performance?	☐	☒	☐
6.29	Are production critical use instruments calibrated regularly?	☒	☐	☐
6.30	Is rework allowed?	☒	☐	☐

SECTION 6. Quality Assurance and Production				
		Yes	No	Not Applicable
6.31	Is reprocessing allowed?	☒	☐	☐
6.32	Are manufacturing and packaging activities traceable to the equipment, areas, and materials used?	☒	☐	☐
6.33	Are production materials handled and stored in a manner to prevent degradation, contamination and cross-contamination?	☒	☐	☐
6.34	If answering 'not applicable' for any of the above, please elaborate: N/A			
Additional Comments: N/A				

SECTION 7. Laboratory Procedures		☐ N/A for this Site		
		Yes	No	Not Applicable
7.1	Does the site have standard procedures for sample handling/tracking?	☒	☐	☐
7.1a	Does the site have standard procedures for retaining samples?	☒	☐	☐
7.1b	Does the site have standard procedures for re-testing samples?	☒	☐	☐
7.2	Does the site have written and approved specifications and test methods?	☒	☐	☐
7.3	Are laboratory instruments calibrated regularly?	☒	☐	☐
7.4	Is there a standard procedure in place for analytical method development?	☒	☐	☐
7.5	Does the site qualify and/or validate analytical test procedures?	☒	☐	☐
7.6	Does the site perform stability testing on materials and/or products?	☒	☐	☐
7.7	Are retention samples of key raw materials maintained?	☒	☐	☐
7.8	Are standards traceable to their preparation and reagents used?	☒	☐	☐
7.9	Are retention samples of finished product maintained?	☒	☐	☐
7.10	Are shelf life/retest/expiration dates available and standardized?	☒	☐	☐
7.11	Does the company provide a certificate of analysis (CoA) and/or a Certificate of Conformation/Compliance (CoC) for each lot or batch?	☒	☐	☐

SECTION 7. Laboratory Procedures		☐ N/A for this Site		
		Yes	**No**	**Not Applicable**
7.12	Does the CoA/CoC contain the manufacture name and location?	☒	☐	☐
7.13	Does the CoA/CoC signed/e-signed by a Quality representative?	☒	☐	☐
7.14	If a repacker performs analyses, will the CoA reflect both the original manufacturing site data as well as the repacking site data?	☐	☐	☒
7.15	If answering 'not applicable' for any of the above, please elaborate: Our material is not sent to a repacker.			
7.16	Additional Comments: N/A			

SECTION 8. Packaging, Storage, and Transport		☐ N/A for this Site		
		Yes	**No**	**Not Applicable**
8.1	Does the site have a validated or qualified labeling system?	☐	☒	☐
8.2	Are batch production records retained and available?	☒	☐	☐
8.3	Are packaging and labeling areas separate from production?	☒	☐	☐
8.4	Are barcode readers in use and challenged regularly?	☐	☒	☐
8.5	Are vision systems in use?	☒	☐	☐
8.6	Is product ever packaged without a label being initially applied (i.e. bright stocking)?	☐	☒	☐
8.7	Do labels include shelf life/expiration dates?	☒	☐	☐
8.8	Do labels include lot/batch number?	☒	☐	☐
8.9	Do labels include requirements for storage conditions?	☒	☐	☐
8.10	Is tamper evident seal used for each container of supplied materials?	☒	☐	☐
8.11	Does the company use a First-In-First-Out or First-Expiration-First-Out system?	☒	☐	☐
8.12	Does the company maintain appropriate storage conditions?	☒	☐	☐
8.12a	Are those storage conditions monitored and documented?	☒	☐	☐
8.13	Does the site make available a description of storage and/or warehouse conditions?	☒	☐	☐
8.14	Does the site distribute products via a third party?	☐	☒	☐
8.15	Are good distribution policies implemented?	☐	☐	☒

SECTION 8. Packaging, Storage, and Transport		☐ N/A for this Site		
		Yes	**No**	**Not Applicable**
8.16	Are transport mechanisms dedicated?	☐	☒	☐
8.17	Does the company validate shipping method?	☐	☒	☐
8.18	Does the company validate packaging methods?	☐	☒	☐
Additional Comments: 8.17 and 8.18, Shipping and packaging methods can be validated if requested by customer.				

I (Supplier) confirm that the information provided in this questionnaire is correct and can be verified.

Date: 21st September 2023

Title:Quality Director