GAEBA Medical Standards – 2024

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GAEBA MEDICAL STANDARDS

**A1.000 Introduction and Purpose**

These standards have been developed to assure consistently acceptable levels of quality, proficiency, and ethics in dealing with ocular tissue for transplantation and define the minimum standards of practice for eye bank functions, as determined by the ophthalmological medical community.

**A1.100 Scope**

These standards are intended to apply to any and all of the eye bank functions, to include:

• Recovery

• Processing

• Storage

• Tissue evaluation

• Donor eligibility determination

• Distribution

These standards shall be reviewed regularly and revised as necessary to incorporate current research findings and improved clinical practice.

**B1.000 Personnel and Governance**

**B1.100 Responsible Party**

All policies and procedures of each eye bank shall be under the supervision of a Responsible Party (RP) appointed by the eye bank’s Board of Directors, Board of Regents or other governing body. The RP shall be responsible for all administrative operations including compliance with all standards.

The RP shall be the individual responsible for the day-to-day operation of the eye bank. It is this individual’s responsibility to carry out policies of the eye bank will be performed by the eye bank, and to prescribe clinically acceptable means to perform these functions.

The RP shall consult with the Medical Director, as well as other medical and legal authorities, in carrying out prescribed responsibilities as necessary. These consultations shall be documented and made available for review during a site inspection.

The RP shall provide all staff members with adequate information to perform their duties safely and competently. Delegation of responsibility for the clinical work of the eye bank shall be as follows:

**B1.200 Medical Director**

The eye bank must have a Medical Director. When the Medical Director is not available, a back-up Medical Director shall be designated who is capable of fulfilling the responsibilities of the Medical Director on an interim basis.

The Medical Director and a back-up Medical Director must be a licensed ophthalmologist who has demonstrated expertise in external eye disease, corneal surgery, research or teaching in cornea and/or external disease.

The Medical Director shall oversee and provide advice on all medical aspects of the eye bank operations. These include but are not limited to:

1. Formulation, approval, and implementation of medical policies and procedures.

2. Participation in training and oversight of technical staff with regard to eye bank functions.

3. Participation in establishment and operation of a quality assurance program.

An eye bank’s Medical Director must observe the designated staff trainer or trainers performing the following procedures as applicable on an annual basis:

1. In-situ corneoscleral disc excision or laboratory corneoscleral disc removal from whole eye

2. Posterior lamellar processing procedure that utilizes a microkeratome

3. At least one type of laser-shaped processing procedure

4. Each manual dissection processing procedure(s) for EK and ALK (i.e., DSEK or DMEK)

If an eye bank Medical Director has not designated any individuals as staff trainers, he/she must observe each technician they have qualified to perform any of the above procedures on an annual basis.

The Medical Director may delegate responsibility for eye bank functions to qualified eye bank personnel; however, the Medical Director shall ensure that the eye bank operates in compliance with all relevant standards. Ultimate responsibility for the suitability of each tissue for the transplantation in patients rests with the transplanting eye surgeon.

An eye bank must replace a Medical Director who has resigned in a timely manner.

**B1.300 Staff Performing Eye Banking Functions.**

The RP may appoint technical and supportive staff and ensure that this staff has the appropriate qualifications and training for the performance of their job responsibilities. The Director shall ensure that there are a sufficient number of qualified eye bank technicians and supportive staff to perform all eye bank laboratory tests and procedures at a level of proficiency established by the bank. The eye bank Medical Director or Medical Director’s designee must document in writing those eye bank tasks in which each staff member is qualified and released to perform independently. C2.000 Training, Certification and Competency Reviews of Personnel Performing Tasks Overseen and/or Regulated by local governing bodies or agencies.

An eye bank or other establishment performing eye banking functions must provide a formal orientation program for each new employee and the employee’s participation must be documented.

An eye bank or other establishment performing eye banking functions, must also establish a comprehensive and well-defined training program outlining specific job-related tasks that each employee is being trained to perform. This comprehensive training program shall include the implementation and documentation of annual competency reviews of the skills and job-related knowledge of all eye bank employees performing eye banking functions. Determination of competency for eye banking functions is the responsibility of the Medical Director or trainer(s) designated and determined competent by the Medical Director. This training program shall contain documentation indicating when each employee is released to perform their job-related tasks independently.

**B2.000 Facilities**

Each eye bank must have sufficient space, equipment and supplies to accommodate the volume of services performed with optimal accuracy, efficiency, sterility, timeliness, and safety.

**B2.100 Eye Bank Laboratory**

The laboratory must be a separate area with limited access in which activities directly related to eye banking are carried out. The laboratory shall have a sink with a drain and running water. There must be adequate counter space for preparation of donor material. The room including walls, floor and sink must be kept clean at all times. Appropriate documentation of regular laboratory cleaning schedules must be maintained and kept on file for a minimum of three years.

Each eye bank laboratory must have an adequate stable electrical source and a sufficient number of grounded outlets for operating laboratory equipment.

**B2.200 Equipment, Maintenance and Cleaning**

Each eye bank laboratory shall have a refrigerator with a device, visible without opening the refrigerator, for recording temperature variations. The temperature recording device should reflect the temperature of the stored tissue under normal storage conditions. Temperature variations must be recorded daily and remain within the range of 2 to 8o Celsius. The refrigerator’s continuous temperature recorder must be calibrated against a standard thermometer as defined by each countries’ regulatory agencies at least once a year. The refrigerator shall be maintained for the use of tissue and tissue preservation solution and must contain clearly defined and labeled areas for all tissue stored, i.e., quarantined tissue, surgical tissue awaiting distribution, and research tissue. Eye banks must detail required refrigerator cleaning intervals and documentation in their Policies and Procedures manual.

In the event of a temperature deviation outside the acceptable range, there must be provision for immediate notification and action to be taken. Testing of the alarm system must be performed and documented in accordance with manufacturer recommendations, but at least annually. The eye bank laminar airflow cabinet or flow hood must be cleaned before and after each use and at regularly scheduled intervals to prevent cross contamination.

Appropriate maintenance and qualification and/or certification records must be maintained on each piece of equipment. These records must show dates of inspection, performance evaluations and any maintenance procedures or repairs performed.

The eye bank must include in its procedures manual, the monitoring, inspection, and cleaning procedures and schedules for each piece of equipment and laboratory area. Documented cleaning schedules for laboratory equipment must be kept on file for a minimum of ten years.

**B2.300 Instruments and Reagents**

**B2.310 Instruments**

To prevent contamination or cross-contamination, all surgical instruments that come in contact with eye tissue during recovery or processing, shall be properly cleaned, decontaminated, and sterilized prior to use and between different donors. The eye bank shall describe in their policy and procedures manual how these activities are performed and monitored. Adequate sterile instruments shall be available for sterile removal and processing of whole eyes and corneas. Instruments shall be inspected frequently to ensure that they function properly. The eye bank’s policy and procedures shall also describe how these instruments are documented and tracked.

An eye bank that sterilizes its instruments shall adhere to local sterilization standards consistent with operating room asepsis policies. The eye bank shall have documentation that show that each load was properly sterilized. The sterilizer shall be

appropriately qualified, and the sterilization process validated before use. The eye bank shall establish and define in their procedures the appropriate sterilization acceptance criteria. The calibration and preventive maintenance routine schedule shall be included in the eye bank’s procedure manual.

If instruments are sterilized by a third party, the eye bank shall qualify the contractor performing the sterilization. The eye bank shall ensure through external audits, that the contractor’s sterilization policies and procedures follow standards and applicable regulations.

The eye bank shall ensure that the qualified sterilizer is routinely calibrated and that the preventive maintenance activities are performed and documented according to manufacturer’s recommendation. If the preventive maintenance is not defined by the manufacturer, then it should be performed annually. Sterilization records shall be maintained and readily available for a minimum of 3 years.

Prior to use, instruments shall be verified to be sterile, that the packaging integrity is intact, the applicable indicators (when available) are acceptable, and that the sterilization expiration dates, if applicable, are acceptable at the time of use.

**B2.320 Supplies and Reagents**

Each eye bank shall have a policy and procedure in which supplies, and reagents used during tissue recovery, processing, or preservation, are properly identified, documented, and tracked. The critical supply/reagent receipt verification shall also be described in the eye bank’s procedure manual.

The supplies and reagents shall be stored and maintained in accordance with manufacturer’s recommendations. Expired supplies and reagents shall not be used for recovery, processing, and preservation of tissue intended for surgical purposes.

The supplies and reagents used during the tissue recovery, processing, or preservation of the ocular tissue must be sterile, where appropriate, and inspected prior to use.

**B2.400 Procedures Manual**

Each eye bank shall maintain its own policies and procedures manual that details all aspects of its specific eye bank functions, and quality assurance

practices. Each procedure must be initially approved, signed, and dated by the Director and Medical Director. An annual review of each eye bank’s procedure with signing and dating by the Director and Medical Director is required. Each eye bank must maintain copies of each procedure it uses and the length of time the procedure was in use. Procedures must be readily available to personnel in the area where operations are performed.

**B2.500 Other Establishments Performing Eye Banking Functions**

All establishments performing specialized or specific eye banking functions must have a Medical Director or access to a Medical Director through a documented consultative relationship.

**B2.600 Infection Control and Personnel Safety**

Written safety procedures for the eye bank operation shall be established in compliance with the relevant regulatory agencies. All eye bank personnel performing functions which may cause exposure to human body fluids or tissue must operate under universal precautions for handling potentially infectious material as determined by the relevant regulatory agency. These written procedures must be included in the eye bank’s procedure manual.

**B2.700 Biohazardous Waste Disposal**

Human tissue and waste items shall be disposed of in such a manner as to minimize any hazard to eye bank personnel and the environment and to comply with the relevant regulatory agency. Dignified and proper disposal procedures shall be used to obviate recognizable human remains and must be documented.

**C1.000 Donor Eligibility Determination**

Before tissue is made available for distribution, the Donor Eligibility Determination must be made by a responsible person. Reference Appendix V for requirements related to the donor eligibility process.

Prior to making an eligibility determination, the donor must be screened according to D1.200. In addition to donor eligibility determination, tissue must be evaluated for suitability per F1.000.

All donors must be identified by name. All prospective donors shall undergo a physical examination as close as possible to the donation with special attention to physical signs of HIV disease, infectious hepatitis, and injecting drug use. Each eye bank shall have a consistent policy for conducting and documenting this examination. Each eye bank shall also have a consistent policy for examination and documentation of the prospective donor’s available medical record and death investigation. Review of all

available records on each donor shall be performed by an individual who is qualified by profession, education, or training to do so, and who is familiar with the intended use of the tissue.

Medical and social histories are important aspects of donor eligibility. Adequately determining eligibility includes, but is not limited to:

1. Infectious disease testing (see D1.200)

2. Physical assessment of the donor (see above paragraph)

3. Tissue evaluation (see F1.000)

4. Donor history evaluation: this must include the donor’s name, social history and donor information obtained from at least one of the following:

a. Pathologist or medical examiner physical assessment of death report

b. Police investigation report accompanied by (a) and/or (c)

c. Medical examiner’s investigative report

d. Donor risk assessment interview

e. Medical record or hospital chart

f. Treating physician interview

g. Medical director oversight to review any donor information where questions

arise in the above areas (see C1.200). This shall be documented.

**C1.100 Donor Screening**

The eye donor’s relevant medical records must be reviewed for:

• specific contraindications (Ref. D1.110): and

• Relevant communicable disease agents and diseases as applicable (Ref. Appendix V): and

• Other diseases as required by the country of import, if exported outside of the country of origin

**C1.110 Contraindications to Transplant**

Determination of donor eligibility is an eye banking function including considerations listed in multiple sources (e.g., U.S. Food and Drug Administration, Health Canada, various state departments, Medical Director input, etc.). In addition to these sources, Tissues from persons with the following are potentially health threatening for the recipient(s) or pose a risk to the success of the surgery and shall not be offered for surgical purposes

recipient(s) or pose a risk to the success of the surgery and shall not be offered for surgical purposes:

A. All Ocular Donors

1. death of unknown cause and there is likelihood of other exclusionary criteria;

2. congenital rubella;

3. Reye syndrome within the past three months;

4. Active viral encephalitis of unknown origin or progressive encephalopathy (e.g., subacute sclerosing panencephalitis, progressive multifocal leukoencephalopathy, etc.);

5. active bacterial or viral meningitis;

6. active bacterial or fungal endocarditis;

7. suspected rabies and persons who, within the past six months, were bitten by an animal suspected to be infected with rabies;

8. intrinsic eye disease;

a. retinoblastoma;

b. malignant tumors of the anterior ocular segment or known adenocarcinoma in the eye of primary or metastatic origin;

c. active ocular or intraocular inflammation: conjunctivitis, keratitis, scleritis, iritis, uveitis, vitreitis, choroiditis, or retinitis;

9. active leukemias;

10. active disseminated lymphomas;

11. Parkinson, amyotrophic lateral sclerosis, multiple sclerosis, and Alzheimer disease;

12. Creutzfeldt-Jakob disease (CJD), variant Creutzfeldt-Jakob Disease (vCJD), or family member with CJD;

13. history of Ebola Virus Disease (EVD);

14. active Lyme disease (known or suspected)

15. history of melanoma with known metastatic disease\*

\* Excluded from this contraindication are tissues subjected to terminal sterilization methods that deactivate neoplastic cells (e.g., gamma irradiation of corneas preserved in albumin).

**C1.120 Screening for Relevant Communicable Disease Agents and Diseases**

Eye banks must screen for relevant communicable disease agents and diseases according to applicable regulations.

**C1.200 Donor Testing**

The eye donor must be tested according to:

• testing requirements (D1.210)

• All required tests in accordance with national and local regulations in the jurisdiction in which they are located

• State requirements, if applicable

• Other testing requirements of the country of import, if exported outside of the country of origin

A review of written results of infectious disease testing shall be received by the eye bank prior to releasing tissue designated for surgical use.

The infectious disease testing laboratory and test kits used must meet relevant regulatory requirements The eye bank must use a laboratory that is accredited by, and whose tests are approved by their own countries’ regulatory agencies.

If plasma dilution sufficient to affect the results of communicable disease testing is suspected, the donor should be considered ineligible, unless a pre- transfusion or infusion sample drawn up to 7 days before recovery is tested; or an algorithm designed to evaluate volumes administered in the 48 hours before specimen collection is used, showing that plasma dilution sufficient to affect the results has not occurred (Ref. Appendix IV).

Eye banks shall use a plasma dilution algorithm which meets the requirements of their own countries’ regulatory agencies. If no such requirements exist, they shall use an algorithm which meets FDA requirements.

**C1.210 Testing Requirements**

The results of the following required testing must be negative or non-reactive for the tissue to be eligible for transplant:

• anti-HIV-1, anti-HIV-2 (or combination test)

• Hepatitis B surface antigen (HBsAg)

• anti-HCV

**C1.220 Non-Required Testing Results**

All non-required positive infectious disease tests must be reported to the eye bank’s medical director, who must review and act on them, or the eye bank must have a standard policy regarding the action to be taken in response to the individual test.

**C1.300 Documentation of Donor Information**

Donor screening forms and/or copies of relevant medical records reviewed must be completed and retained on all donated eye tissue as part of the donor record. See Section L1.000.

A unique donor identifying number, i.e., medical examiner or coroner case number, hospital medical record number, social security or driver’s license number, shall be obtained and recorded in the donor record.

**C1.400 Method and Authorization for Donation**

Documentation of legal authorization for recovery is essential for medical-legal reasons. Authorization procedures and forms must conform with relevant regulatory requirements.

**C1.500 Donor Age**

Since no definite relationship has been established between the quality of donor tissue and age, the upper and lower age limit is left to the discretion of the Medical Director.

**C1.600 Interval between Death, Enucleation, Excision, Preservation, and Processing**

Acceptable time intervals from death, enucleation or excision to preservation may vary according to the circumstances of death and interim means of storage of the body. It is generally recommended that corneal preservation occur as soon as possible after death. All time intervals for each donor, i.e., the time of death to the time of enucleation and preservation and/or the time to corneal excision, and/or the time to additional tissue processing, shall be recorded. The time that cooling of ocular tissues and/or refrigeration of the body was begun shall be recorded, if applicable.

**C1.700 Eye Maintenance Prior to Recovery**

The prospective donor’s corneal integrity should be maintained. Each individual eye bank’s procedure is left to the discretion of the Medical Director and shall be clearly documented.

**C1.800 Living Donors**

Eye tissue that is removed and intended for surgical use from a living donor shall have the same standards applied as for all cadaveric tissue, e.g., the same donor medical history shall be obtained, the same records, infectious disease tests, etc. No extended quarantine period, outside the usual 24-48 hours for infectious disease test results, shall be required for corneal tissue used for transplantation that is stored in short or intermediate term storage solution.

**D1.000 Recovery, Processing, and Preservation**

Recovery, processing, and preservation must be done using aseptic technique as defined by relevant standards in each country or state

All procedures must be documented.

The Medical Director is responsible for establishing the eye bank’s procedures for recovery, processing, and preservation of tissue. The Medical Director and Director are responsible for assuring that eye bank personnel comply with all applicable procedures for the recovery, processing, and preservation of tissue.

**D1.100 Recovery**

The donor’s identity shall be verified prior to recovery. Recovery may be performed via enucleation or in situ method.

A 5% povidone-iodine (PI) solution shall contact the entire surface of any ocular tissue intended for transplantation at least twice between the time of the donor’s death and tissue preservation (e.g. corneoscleral disc in corneal preservation solution or whole eye in moist chamber). Regardless of how PI is administered (e.g., # of drops, a specific mL, soak, etc.), the amount must be sufficient to completely cover the corneal surface, conjunctiva, lids, and lashes. The contact time for each application should not be less than 2 minutes and not exceed 5 minutes. Povidone-iodine solution should be irrigated from the ocular surface with a sterile eye wash/irrigating solution between applications and prior to preservation. This concentration, volume of solution, and the duration of ocular surface exposure to the solution shall be specified in the eye bank’s operating procedures.

The corneoscleral disc shall initially be examined with a penlight or portable slit lamp for clarity, epithelial defects, foreign objects, contamination, and scleral color prior to enucleation or in situ corneoscleral disc excision.

Standard Precautions shall be followed during donor physical examination, recovery, and all tissue handling procedures to protect eye bank staff from potential exposure to infectious diseases. Tissue from donors with the following is hazardous to eye bank personnel:

• Active Viral Hepatitis

• Acquired Immunodeficiency Syndrome (AIDS) or HIV seropositivity

• Active viral encephalitis or encephalitis of unknown origin

• Creutzfeldt-Jakob Disease (CJD)

• Rabies

**D1.200 Processing and Preservation**

Aseptic processing shall be performed in a certified and qualified bacteriologically and climate-controlled environment. The processing shall be performed by qualified, trained, and competent staff using validated methods to prevent contamination and cross-contamination and to maintain tissue quality for its intended use.

Eye banks shall establish and maintain policies, processes, and procedures designed to minimize contamination of the tissue. The following shall be addressed:

1. Environmental controls and monitoring

2. Process controls

3. Staff training in aseptic technique

4. Attire, gowning, and use of personal protective equipment

5. Workflow movement and movement of personnel through workspaces

6. Prevention of cross-contamination and labeling mix-ups (e.g., tissue from different donors may not be processed simultaneously.

**D1.210 Whole Eye**

Eye banks that preserve and store whole eyes for lamellar or refractive keratoplasty may use preservation methods such as moist chamber at 2-8 degrees Celsius, freezing below zero degrees Celsius, or some other validated method.

**D1.220 Cornea**

Eye banks that process corneas intended for transplant may use one of the following methods. The Medical Director must develop appropriate tissue selection criteria and approve the procedure for each method utilized. All processes must be validated. When changes or process deviations occur, the eye bank shall review and evaluate the process and perform revalidation where appropriate.

**D1.221 Processing via Excision of the Corneoscleral Disc from Enucleated Whole Eyes**

**D1.222 Lamellar Tissue Processing**

Processing of lamellar tissue may be performed using manual or automated methods (e.g., microkeratome).

**D1.223 Laser Assisted Processing**

Lasers may be used to process lamellar tissue or custom wound architecture (e.g., femtosecond laser).

**D1.230 Sclera**

There are various methods of processing sclera, including utilizing

ethanol (70% or greater ethyl alcohol), sterile glycerol, cryopreservation, radiation sterilization, or some other validated method.

**D1.300 Use of Short or Intermediate Term Storage Solution**

Eye banks shall use an appropriate corneal storage solution. The solution shall be used and stored according to the manufacturer’s recommendations for temperature, date, and other factors. The manufactured solution purchased and shipped to the eye bank shall be inspected for damage upon arrival. The lot number of storage solution used for each cornea shall be recorded on the tissue report to allow tracking and recall.

**D1.400 Long Term Preservation and Organ Culture**

Some eye banks employ preservation techniques such as long-term preservation of corneal tissue and organ culturing. An eye bank that uses long term preservation and/or organ culture shall carefully document the procedure(s) in their procedures manual, and adhere to rigid aseptic technique.

**E1.000 Tissue Evaluation**

The performing of any and /or all of the following on corneal tissue intended for transplant: slit lamp examination, endothelial cell density, and pachymetry measurement; according to Matrix I.

The ultimate responsibility for determining the suitability of the tissue for transplantation rests with the transplanting surgeon.

The following evaluations must be performed prior to deeming preserved and/or processed tissue suitable for intended use.

|  |  |
| --- | --- |
|  | **Matrix I: Tissue Evaluation Requirements** |
| **Suitability for** | **Slit Lamp Biomicroscopy** | **Specular Microscopy** | **Pachymetry** |
| **Preserved Ocular Tissue – Not Processed by Eye Bank** | PKP | Required | Required | Not required |
| DSAEK/DSEK | Required | Required | Not required |
| DMEK/DMAEK/ PDEK | Required | Required | Not required |
| ALK | Required | Not required | Not required |
| Long-term preservation (whole or sectionedcornea or sclera) | Required2 | Not required | Not required |
| **Eye Bank Processed Ocular Tissue** | PKP (laser-processed) | Required | Required1 | Not required |
| DSAEK | Required | Required1 | Required (graft thickness) |
| DMEK/PDEK | Required | Required1 | Not required |
| DMAEK | Required | Required1 | Required |
| ALK | Required | Not required | Required (graft thickness) |
| Long-term preserved (whole or sectionedcornea or sclera) | Required2 | Not required | Not required |
| 1 In lieu of specular microscopy, a validated method for assessment of endothelium after processing meets this requirement. |
| 2 In lieu of slit lamp biomicroscopy, gross examination to ensure graft is free of contamination meets this requirement. |

**E1.100 Slit Lamp Examination**

The corneoscleral disc shall be examined for epithelial and stromal pathology and in particular endothelial disease using slit lamp biomicroscopy. Whole eyes to be distributed for lamellar processing must have the same examination. Corneoscleral discs that have been processed for lamellar keratoplasty procedures shall be re-evaluated by slit lamp biomicroscopy to ensure that there was no damage to the relevant transplantable tissue.

Document the observations of the slit lamp examination with particular attention to the epithelium, stroma, and endothelium such as, but not limited

to, scars, edema, arcus, pterygia, neovascularization, striae, epithelial defects, guttata, polymegathism, pleomorphism, infiltrates, or foreign bodies.

The eye bank should delineate a “clear zone” on each cornea as a measurement of diameter (in millimeters) of the clear central cornea, free of neovascularization, pterygia, arcus, or other stromal anomalies. Anterior stromal scars may be omitted from clear zone measurement, as long as details of scar location, size, and relative depth are made available to the transplanting surgeon. Clear zone measurements are acknowledged to impact surgical suitability determination more significantly for surgery types utilizing the anterior corneal segment (e.g., PK, ALK, K-Pro). Eye banks are encouraged to provide a measurement of an arcus clear zone (a measurement of clear central cornea free of arcus only) if that measurement may responsibly improve or otherwise clarify surgical suitability determination.

**E1.200 Endothelial Cell Density and Pachymetry**

Determination of endothelial cell density via specular microscopy (or quantitative light microscopy for organ cultured corneas) shall be a standard method of corneal tissue evaluation (according to Matrix I). Minimal endothelial cell count limits are left to the discretion of the Medical Director. When it is impossible to obtain an endothelial cell count, this requirement may be waived on a case-by-case basis by the Medical Director. Calibration of endothelial cell counting equipment shall be done according to manufacturer guidelines, when applicable, and on at least an annual basis. Calibration procedures shall include specific directions and limits for accuracy.

**E1.300 Determination of Surgical Suitability**

The eye bank responsible for evaluation of ocular tissue shall specify whether the tissue meets the criteria for penetrating keratoplasty (PK), anterior lamellar keratoplasty (ALK/DALK), Descemet’s stripping endothelial keratoplasty (DSEK/DSAEK), Descemet’s membrane endothelial keratoplasty (DMEK), Pre-Descemet’s endothelial keratoplasty (PDEK), keratolimbal allograft, and “other” surgical use (e.g., keratoprosthesis, long-term preservation for later shunt patch/ALK/tectonic use, experimental surgical use, etc.).

Corneoscleral Disc Minimum Suitability Standards

Minimum suitability for penetrating keratoplasty (PK):

• No infiltrates

• No pterygia, neovascularization, foreign bodies, or visually significant stromal scars within intended graft area

• No Descemet’s membrane detachment or tears within intended graft area

• No evidence of endothelial dystrophy

• Minimum endothelial cell density (as defined in eye bank’s policy)

• No Down syndrome or evidence of ectatic dystrophy (e.g., keratoconus, keratoglobus, etc.,

• No prior laser or incisional refractive surgery (e.g., radial keratotomy, lamellar inserts, photoablation, etc.)

Minimum suitability for anterior lamellar keratoplasty (ALK/DALK):

• No infiltrates

• No pterygia, neovascularization, foreign bodies, or visually significant stromal scars within intended graft area

• No Down syndrome or evidence of ectatic dystrophy (e.g., keratoconus, keratoglobus, etc.)

• No prior laser or incisional refractive surgery (e.g., radial keratotomy, lamellar inserts, photoablation, etc.)

Minimum suitability for Descemet’s stripping endothelial keratoplasty (DSEK/DSAEK):

• No infiltrates

• No foreign bodies or visually significant scars affecting posterior stroma within intended graft area

• No Descemet’s membrane detachment or tears within intended graft area

• Minimum endothelial cell density as defined in eye bank’s policy

• Sufficient rim size and corneoscleral disc size to facilitate mounting on artificial anterior chamber

Minimum suitability for Descemet’s membrane endothelial keratoplasty (DMEK), Pre-Descemet's endothelial keratoplasty and Descemet's membrane automated endothelial keratoplasty. (DMEK/PDEK/DMAEK):

• No infiltrates

• No foreign bodies

• No Descemet’s membrane tears within intended graft area

• Minimum endothelial cell density as defined in eye bank’s policy)

Minimum suitability for keratolimbal allograft (KLA):

• No infiltrates

• Sufficient scleral rim (minimum must be defined in eye bank’s policy)

• Conjunctiva must be intact over sufficient portion of rim to facilitate allograft (rim portions may be considered from mated pairs)

• No history of melanoma or metastatic cancer of a solid organ

Minimum suitability for Keratoprosthesis (K-Pro):

• No infiltrates

• No pterygia, neovascularization, foreign bodies, or significant corneal thinning

• No prior refractive surgery (e.g., radial keratotomy, lamellar inserts, photoablation, etc.)

• No Down syndrome or evidence of ectatic dystrophy (e.g., keratoconus, keratoglobus, etc.)

Minimum suitability for Long-Term Cornea Preservation/Other:

• No infiltrates

• No pterygia on graft segments

Sclera Minimum Suitability Standards

Minimum suitability for sclera for any surgical use:

• No infiltrates on cornea from the eye that produced scleral grafts

• No history of melanoma or metastatic cancer of a solid organ

**F1.000 Quality Assurance**

Each eye bank shall have a formally established quality assurance program. This program shall include:

• Establishment and maintenance of procedures for all functions performed by the eye bank (including review, approval, and revision)

• Monitoring and evaluation of functions through periodic audits by an individual(s) not regularly involved in the processes being monitored

• Identification of problems and complaints relating to activities (receiving, investigating, evaluating, and documenting information relating to eye banking requirements)

• Development of plans for corrective actions, including monitoring for effectiveness

The quality assurance program shall address applicable requirements relating to the following areas:

1. Facilities

2. Environmental control

3. Equipment

4. Supplies and reagents

5. Recovery

6. Processing and processing controls

7. Labeling controls

8. Storage

9. Receipt, pre-distribution shipment, and distribution

10. Donor eligibility determinations, donor screening, and donor testing

11. Tissue evaluation

Each eye bank shall document all aspects of its quality assurance program. Records relating to the quality assurance program shall be maintained for a minimum of ten years or a time frame according to relevant regulatory standards. These records shall be made available for any relevant inspections.

The Quality Assurance Program shall establish a system for reporting, documenting, and investigation of deviations. Deviations for distributed tissue relating to eye bank functions must be reported in a manner according to relevant regulatory standards.

The eye bank’s quality assurance program shall include a method for the receiving surgeon to report adverse reactions from the transplantation of corneal, scleral, or other ocular tissue to the distributing eye bank. The distributing eye bank must forward the adverse reaction information to the source eye bank, which made the donor eligibility determination. The source bank must perform an investigation and must report the adverse reaction information in a manner according to relevant regulatory standards.

The source bank must notify all entities involved in the recovery, processing, storage, distribution, tissue evaluation, and donor eligibility determination of the results of the investigation. Each of the involved entities must maintain documentation of the adverse event and results of the investigation forwarded to it by the source bank.

Infection of a systemic nature that the medical director’s investigation determines to be possibly, likely/probable or definitely due to donor tissue must be communicated to all entities that recovered organs or received or recovered tissues from that donor.

The Medical Director shall receive and review all adverse reaction reports, documenting any corrective actions he/she determines are indicated.

**F1.100 Quality Control**

The Director shall prescribe tests and procedures for measuring, assaying or monitoring properties of tissues essential to the evaluation of their safety for transplantation, e.g., hepatitis B surface antigen and human immunodeficiency virus (HIV) antibody, and conform with relevant regualtory. Results of all such tests or procedures, together with evaluations based on these findings, shall become part of permanent record of all tissues intended for surgical use.

**F1.200 Microbiologic Culturing**

Culturing of eye bank donor eyes may be performed despite the recognition by many that bacteriologic contamination of donor eyes does not necessarily lead to infection and that presurgical or surgical cultures may not correlate with postoperative infection if it should occur. Cultures may be performed either before and/or at the time of surgery.

a. Presurgical Cultures

Eye banks may elect to perform corneoscleral disc cultures at the time of corneal preservation in storage solution. Positive culture reports shall be reported to the receiving surgeon or recipient eye bank.

b. Surgical Culturing

Each eye bank shall indicate on the information sheet accompanying the tissue for transplantation whether corneoscleral disc cultures were performed prior to distribution. Positive results in cases of postoperative infection shall be reported to the eye bank that recovered the tissue as well as to the eye bank that distributed the tissue.

**F1.300 Tissue Recall or Tissue Withdrawal**

Eye banks must have a policy and procedure for potential recall of tissue.

Positive test results or information about behavioral risks or medical history, received after release of tissue, that indicate a risk for transmission of a relevant communicable disease must be reported to the:

• Eye bank's medical director

• Consignee (i.e., the transplanting surgeon, processor or distributing eye bank)

• Appropriate government agency, within the relevant timeframe

**G1.000 Non-Surgical Donor Tissue**

The use of ocular tissue from a donor determined to be ineligible is not prohibited for non-clinical uses, so long as they bear the Bio-hazard legend and are labeled “For Non-Clinical Use Only” and “Not for Transplant.”

Tissue being stored for non-clinical purposes (e.g., teaching and/or research) from a donor who has been determined to be ineligible for transplantation due to results of required testing and/or screening or from donors who have not been tested for required infectious diseases, must have a label affixed to the individual tissue container which contains the information below.

1. “For Non-clinical Use Only”

2. “Biohazard” or Biohazard legend

3. “Not for Transplant”

A label containing the required information listed above must be affixed to the individual tissue container for tissue that is distributed for non-clinical purposes.

**H1.000 Storage**

All tissue shall be transported and stored in quarantine from the time of recovery until the donor eligibility determination has been completed. Quarantined tissue must have a label designating the tissue as “quarantine” affixed to the individual tissue container.

All surgical tissue shall be stored in quarantine in a physically separate area clearly identified for such use, or through use of other procedures such as automatic designation, until a donor eligibility determination has been made.

If a donor is determined ineligible, the tissue must be stored or identified in a physically separate area labeled for such use, or other procedures must be followed that are adequate to prevent improper release.

All tissue shall be stored aseptically at a temperature appropriate to the method of preservation used. Eye banks must precisely document their procedures for storage of corneal tissue, whether it is in the form of the whole eye or the corneoscleral disc in an appropriate storage solution.

**H1.100 Expiration Dating**

Where appropriate, an expiration date must be assigned based upon methods of processing, preservation, storage, and packaging.

**I1.000 Labeling**

All ocular tissue distributed for surgical use shall be in a container which is clearly and indelibly labeled to include at least the information below.

1. Name of source eye bank.

2. Tissue identification number.

3. Type of tissue (e.g., cornea, whole eye, sclera).

4. If cornea has had additional processing (e.g., lamellar, laser shaped), clearly indicate this on the label.

6. Expiration date of tissue, in the international format (YYYY-MM-DD).

7. A statement that the tissue is not to be considered sterile unless tissue has been subjected to a validated process to ensure sterility.

8. Type of storage solution.

**J1.000 Distribution of Tissue**

**J1.100 Review of Donor Medical Information**

Prior to distribution of tissue for transplantation, the Medical Director or his/her designee shall review and document that the medical and laboratory information is in accordance with medical standards, and that any departures from procedure do not increase the risk of communicable disease transmission.

**J1.200 Distribution Compliance**

An eye bank performing distribution shall inform the consignee, in writing, of requirements for tracking and traceability, outcomes and adverse reaction reporting. Compliance with applicable laws, regulations and standards in eye bank functions performed after distribution is the responsibility of the consignee.

**J1.300 Fair and Equitable System**

Eye banks shall establish and document a system of distribution for transplant tissue that is just, equitable and fair to all patients served by the eye bank. Documentation of distribution requests for, offers of, and delivery of eye tissue shall be available for inspection by and relevant regulatory agencies. Access to tissue shall be provided without regard to recipient sex, age, religion, race, creed, color, or national origin.

**J1.400 Returned Tissue**

For transplant tissue returned and redistributed, tissue transportation and storage information must be documented and made available to the eye bank and transplanting surgeon for short and intermediate-term preserved transplant tissue. For special research studies, as approved by relevant regulatory agencies, such tissue transportation and storage information may be withheld from the transplanting surgeon.

**J1.500 Fraudulent Activity**

If the eye bank discovers fraudulent activity has occurred in the distribution, shipping or labeling of any tissue imported or exported by the eye bank, an investigation shall be performed to identify the root cause of the occurrence. The eye bank shall report the occurrence and their findings to any relevant regulatory agencies.

**K1.000 Documentation to Accompany Donor Tissue**

**K1.100 Tissue Report Form**

In special circumstances, like approved research programs, certain label and tissue report form requirements may be waived with approval from any relevant regulatory agencies.

Tissue distributed for transplant use shall be accompanied by a tissue report form and may include additional forms to address all requirements. The tissue report, together with pertinent additional forms shall contain the following:

All Tissues:

1. Name of (Source) eye bank

2. Location of eye bank

3. Telephone number of eye bank

5. Type of storage solution

6. All dates shall be written as YYYY-MM-DD HH:MM.

7. If cornea is processed, clearly indicate the type of processing performed or the indicated use (e.g., endothelial keratoplasty, posterior lamellar keratoplasty, anterior lamellar keratoplasty, laser assisted keratoplasty, etc.).

8. Tissue evaluation reporting requirements according to Matrix II.

9. A summary of records reviewed regarding the eligibility of tissue for transplant.

|  |
| --- |
| **Matrix II: Reporting Requirements** |
| **Content on Tissue Report Form** | **Unprocessed Tissue** | **Processed Tissue** |
| **Short or Intermediate-****Term Storage** | **Intermediate-Term Storage\*** | **Long-Term Storage\*\*** |
| Donor age | Required | Required | Not Required |
| Donor cause of death | Required | Required | Not Required |
| Donor death date and time | Required | Required | Not Required |
| Preservation date and time | Required | Required | Not Required |
| Additional processing date and time |  | Required | Not Required |
| Date and time that cooling ofocular tissues or body refrigeration began | Required | Required | Not Required |
| Name/identifier of technician who recovered tissue | Required | Required | Not Required |
| Name/identifier of technician who initially preserved (stored)tissue | Required | Required | Not Required |
| Name/identifier of technician(s) who evaluated tissue | Required | Required | Not Required |
| Name/identifier of technicianwho processed tissue |  | Required | Not Required |
| Morphology and dimensions of processed tissue | Required | Required |
| Diameter of processed graft | Required | Not Required |
| Pachymetry (graft thickness) | Not Required | Required | Not Required |
| Slit lamp observations | Required | Required | Required (othervisual exam acceptable) |
| Specular microscope observations (including endothelial cell density) | Required (unless whole eye, anterior or tectonic useonly) | Required1 (unless anterior or tectonic use only) | Not Required |
| Suitability for indicated surgical uses | Required | Required | Required |

\* Intermediate-Term Storage - For example: manual, microkeratome, or laser-processed cornea for penetrating, endothelial (e.g., DSAEK, DSEK, DMEK, DMAEK), or anterior lamellar keratoplasty.

\*\* Long-term storage - For example: frozen whole globe, cornea in glycerin, irradiated cornea, sclera.

1 In lieu of specular microscopy, a validated method for assessment of endothelium after processing meets this requirement.

**K1.200 Package Insert Form**

A “Package Insert” form shall accompany the tissue for transplantation. This form shall include the following:

1. Recommended storage temperature for specific type of tissue (cornea; sclera; whole eye). Specific emphasis on DO NOT FREEZE for corneas.

2. That the surgeon should check for integrity of the seal and immediately report to the eye bank any evidence of possible tampering.

3. For corneas in intermediate-term storage solution, a color change per the manufacturer’s guidelines may indicate a change in pH, in which case the tissue should not be used, and a report made immediately to the eye bank.

4. Whether pre-surgical microbiologic cultures were performed by the eye bank.

5. The form shall also advise the receiving surgeon that the tissues are delivered with no warranty as to merchantability or fitness for a particular purpose, and that the receiving surgeon is ultimately responsible for judging if the tissue is suitable for use.

6. The form shall advise the consignee that they are responsible for tracking of:

• The tissue recipient’s name and unique identification number,

• Age and/or date or birth, diagnosis, date of surgery, location of surgery, type of surgery,

• The name of the transplanting surgeon when the tissue is transplanted, and

This information is needed to track the tissue from the donor to consignee and from the consignee to the recipient.

7. Infectious disease tests were performed by a valid serology laboratory as defined by country of origin.

8. That tests were used for infectious disease testing as required by the relevant regulatory agencies, some of which are approved for pre-mortem blood and approved tests for cadaveric blood were used where available.

9. A list of infectious disease test results for that specific donor.

This information may be included on the eye bank’s donor screening form as long as it is easily noticed; otherwise, a separate package insert form is advised.

**K2.000 Packaging, Sealing and Packing for Transport**

Each tissue for distribution shipment shall be individually packaged and sealed with a tamper-evident seal or enclosed in a tamper evident container.

Each tissue for predistribution shipment transported by a secondary carrier shall be sealed with a tamper-evident seal or enclosed in a tamper-evident container.

Corneal tissue in intermediate-term storage solution shall be packaged using a method designed to maintain cool conditions and prevent freezing. The package content should demonstrate remaining coolant effect at the time of use or removal to mechanical storage or replacement of the coolant. For tissue preserved by other methods such as long-term preserved, organ culture, or short-term preserved tissue, the packaging method shall be appropriate to the method of preservation used. Packing shall be done so that the tissue label and documentation to accompany the tissue do not become wet. Special instructions shall be included on a Package Insert. See Section L1.200.

**L1.000 Eye Bank Records**

**L1.100 Length of Storage**

All records shall be kept for a time period determined by the relevant regulatory agencies. In the absence of a regulatory standard, it is recommended that the records are retained for a minimum of ten (10) years.

**L1.200 Confidentiality**

All eye bank records and communications between the eye bank and its donors and recipients shall be regarded as confidential and privileged.

**L1.300 Donor Screening Forms**

Donor screening forms shall contain information regarding the circumstances surrounding the death of a donor and adequate medical history so that the eligibility of the tissue for transplantation may be judged.

**L1.400 Minimum Information to Be Retained**

Forms for retaining donor and recipient, distributing eye bank, or consignee information shall be established and shall be readily accessible for inspection as required by the relevant regulatory agencies. Eye bank records shall include the following minimum information:

See Section L1.000 for information to be included on the Tissue Report Form.

1. Name of eye bank

2. Type of storage solution

3. Storage solution lot numbers

4. Unique donor identification number

5. Name of donor (or if import tissue, name of importing eye bank and their unique ID number)

6. Age of donor

7. Cause of death

8. Death date and time

9. Enucleation or in situ excision date and time

10. Preservation date and time

11. The date and time that cooling of ocular tissues and/or refrigeration of the body was begun

12. Additional tissue processing date and time

13. Slit lamp report(s)

14. Endothelial cell density(ies) (if applicable)

15. Unique identifier of enucleator/processor/evaluator/technician. Unique identifier (name, ID #, etc.) shall be clearly cross referenced to staff training, certification, and competency review records and readily available for Accreditation Board members as part of inspection.

16. Name of surgeon, distributing eye bank, or consignee receiving tissue

17. Tissue readily traceable from donor to distributing eye bank or consignee for each tissue identifier (See Section M1.500)

18. Date, time, method of transportation

19. Utilization of tissue: i.e., surgical, research, training

20. Printed or electronic results of all required and non-required infectious disease screening tests

21. Microbiologic screening results, if performed

22. Microbiologic reports of positive donor rim cultures from the receiving surgeon, if reported

23. Adverse reactions, if reported

24. Documentation that post-operative outcome information from the transplanting surgeon has been requested

**L1.500 Recipient Follow-Up Information**

1. Each distributing eye bank shall obtain consignee name and address information for each eye tissue used for human transplantation distributed by the bank.

2. Each distributing eye bank shall seek:

• Patient’s name (if allowed by law)

• Unique recipient identification, such as:

a. Social security number

b. Driver’s license number

c. Medical Record number

d. Alien identification

e. Passport number

f. Other unique identifier appropriate to the health care delivery system where surgery is performed

• Age and/or Date of Birth

• Diagnosis

• Name of surgeon receiving transplanting tissue

• Date of surgery

• Location of surgery

• Post-operative complications (tissue related)

• Type of surgery performed, e.g., penetrating keratoplasty, anterior lamellar keratoplasty, endothelial keratoplasty, keratolimbal allograft, and/or tectonic

3. Corneas and scleral tissue that can be used beyond 14 days post-mortem may be stocked at an institution only if it is for single patient use; the distributing eye bank must be able to track the tissue to the consignee.

4. Each distributing eye bank must request postoperative outcome information between three and six months after transplant from the consignee concerning possible adverse reactions on all cornea tissue, except long term preserved, used for human transplantation that was distributed to the consignee by that bank. This request must be addressed to the transplanting surgeon and delivered separately from the documentation that accompanies the eye tissue. For special research studies where postoperative outcomes are monitored by other means, by recommendation of the relevant regulatory agencies, eye bank solicitation of postoperative outcome information and documentation of such solicitation (under M1.400 item 25) will not be required.

**M1.000 Amendments**

These standards may be amended as medical technology, techniques and information require. A comment period may be provided prior to the intended effective date.

**Glossary**

Definition of Terms

**Ambient Storage.** Stored in a solution at ambient temperature.

**Annually.** A period less than or equal to 12 calendar months.

**Anterior and Posterior Layers.** A pre-cut cornea where both the anterior and posterior layers are present.

**Anterior Lamellar Keratoplasty (ALK).** Transplantation of the anterior stroma of the cornea.

**Anterior Layer.** Corneal stroma without endothelium. May include epithelium.

**Aseptic Technique.** Method by which contamination with microorganisms is prevented.

**Audit.** A documented review of procedures, records, personnel functions, equipment, materials, facilities, and/or vendors to evaluate adherence to the written SOP, standards, or federal, state and/or local laws and regulations.

**Autologous use.** The implantation, transplantation, infusion, or transfer of human cells or tissue back into the individual from whom the cells or tissue were recovered.

**CJD.** An abbreviation for Creutzfeldt-Jakob Disease.

**Colloid.** A protein or polysaccharide solution such as albumin, dextran, or hetastarch that can be used to increase or maintain osmotic (oncotic) pressure in the intravascular compartment, or certain blood components, such as plasma and platelets.

**Complaint.** Any written or oral communication concerning dissatisfaction with the identity, quality, packaging, durability, reliability, safety, effectiveness, or performance of tissue.

**Competency.** The ability of an employee to acceptably perform tasks concomitant with his/her educational level for which he/she has been trained.

**Competency Assessment.** The evaluation of the ability of an employee to acceptably perform tasks that are expected of the employee for the duties/responsibilities assigned to him/her.

**Conjunctiva.** Transparent mucous membrane passing over the inner surface of the eyelids and reflected over the front part of the sclera

**Consent.** A process where approval for donation is obtained from the donor (called “First Person consent” or “FP”) or the donor’s next of kin or other legally recognized representative.

**Consignee.** Any eye bank not accredited by EBAA, third-party distributor or transplanting surgeon, surgical facility, researcher, or educational facility (whether individual, agency, institution, or organization) that receives tissue and assumes responsibility for any step in the processing, storage, distribution and/or use of such tissue.

**Container.** A receptacle that is used to contain tissues and is in direct contact with the tissue.

Contract Services. Those functions pertaining to the recovery, screening, testing, processing, storage, and/or distribution of tissue that another establishment agrees to perform for an eye bank.

**Cornea.** Transparent anterior part of the outer fibrous coat of the eye bounded by an outer stratified epithelium and an inner monolayer of endothelial cells. The major refractive component of the eye.

**Corneal Button.** Cornea with scleral rim removed.

**Corneoscleral Disc.** Cornea excised with scleral rim which may include some conjunctiva.

**Critical Supplies and Reagents.** Supplies or reagents that come in contact or can affect the quality of the ocular tissue during the recovery or processing of the tissue.

**Cross-Contamination.** The transfer of infectious agents from one tissue to another tissue, or from one donor’s tissue to another donor’s tissue.

**Cryopreserved.** Preserved by freezing or vitrification in the presence of a cryoprotectant and using a method validated to maintain cellular viability and/or preserve tissue matrix structure. The information about the cryoprotectant may be specified using the storage solutions attribute group or on the tissue container label or in accompanying documentation.

**Crystalloid.** A balanced salt and/or glucose solution, such as saline, TPN (total parenteral nutrition), Ringer’s lactate solution, or 5 percent dextrose in water, used to replace electrolytes or to increase intravascular volume.

**Decontamination.** The use of physical or chemical means to remove, inactivate, or destroy bloodborne pathogens on a surface or item to the point where they are no longer capable of transmitting infectious particles and the surface or item is rendered safe for handling, use, or disposal.

**Dehydration.** The removal of water from cells and/or tissue.

**Departure from Procedure/Accident.** An intended change from an established procedure, including a standard operating procedure (SOP), which occurs before the HCT/P is distributed and is consistent with applicable regulations and standards.

**Descemet’s Membrane Automated Endothelial Keratoplasty (DMAEK).** Transplantation of a peripheral ring of stroma along with a central area of Descemet’s membrane and endothelium.

**Descemet’s Membrane Endothelial Keratoplasty (DMEK).** Transplantation of endothelium and Descemet membrane only.

**Descemet’s Stripping Automated Endothelial Keratoplasty (DSAEK).** Transplantation of corneal endothelium, Descemet membrane, and posterior stromal layers of varying thickness.

**Designee.** A person authorized by responsible party to perform assigned duties.

**Deviation/Error.** An event that represents a deviation from applicable regulations, standards, or established specifications, or is unexpected or unforeseeable.

**DIN.** An abbreviation for Donation Identification Number.

**Disinfectant.** An agent that reduces the number of viable cellular microorganisms.

**Disinfection.** A process that reduces the number of viable cellular microorganisms, but does not necessarily destroy all microbial forms, such as spores and viruses.

**Disposition.** The final destination of tissue, including use for transplantation, education, research, or discard.

**Distributing Eye Bank.** The last eye bank that provides tissue to a consignee. A process must be in place to ensure the principles of tracking, traceability, and adverse event reporting.

**Distribution.** A process of allocation of tissue for transplant, research, or educational use. This process includes receipt of request, selection, inspection, and release of tissue, to a consignee such as a surgeon, surgical center, or educational research center. The principles of tracking, traceability and adverse reaction reporting will be maintained throughout the process of distribution.

**Donation Identification Number (DIN).** A unique identification of a donation/recovery event.

**Donor.** A living or deceased individual who provides the source of tissue for transplantation, education, or research.

**Donor Eligibility Determination.** The evaluation of all available information about a potential donor to assess whether the donor meets qualifications specified in the SOP and standards. This includes, but is not limited to medical, social, and sexual histories; laboratory test results; physical assessment or physical examination; and autopsy findings (if performed).

**Donor Referral Sources.** Entities such as hospitals, medical examiners, coroners and individual allied health care professionals or others who identify potential eye tissue donors and refer them to eye banks.

**Donor Risk Assessment Interview.** A documented dialogue in person or by telephone with an individual or individuals who would be knowledgeable of the donor’s relevant medical history and social behavior. For example, this may be the donor, if living; the next of kin; the nearest available relative; a member of the donor’s household; other individual with an affinity relationship (e.g., caretaker, friend, significant life partner); and/or the primary treating physician. Alternatively, a living donor may complete a written questionnaire. Relevant social history is elicited by questions regarding certain activities or behaviors that are considered to place such a potential donor at increased risk for a relevant communicable disease agent or disease.

**Donor Screening.** See GAEBA Medical Standard C1.100.

**End User.** A hospital, surgeon, surgicenter, research center or any entity that utilizes tissue provided by an eye bank.

**Endothelial Keratoplasty.** Transplantation of the corneal endothelium in various forms.

**Enucleation.** Recovery of the whole eye.

**Equipment Qualification.** Protocols designed to adequately evaluate, prior to use, whether or not pieces of equipment will perform to expectation, and normally function within the required tolerance limits.

**Evaluation.** The assessment of an entity, tissue, equipment, personnel, performance in relation to predetermined expectations or standards.

**Expiration Date.** The date after which instruments, supplies or tissues are deemed no longer suitable for use.

**Eye Bank.** An entity that provides or performs one or more functions involving ocular tissue from living or deceased individuals for transplantation, research, and/or educational purposes. See Eye Bank Functions.

**Eye Bank Functions.** Functions performed by an eye bank that subject the eye bank to accreditation policies of the EBAA, including 1) recovery, 2) processing, 3) storage, 4) tissue evaluation, 5) donor eligibility determination, and 6) distribution.

**FIN(P).** An abbreviation for Processing Facility Information Code.

**Fraudulent activity.** Any activity that involves potentially falsified tissue, tissue documents or labeling of tissue containers of imported or exported tissue for transplant or research

**Freeze Dried.** (Lyophilized) Preservation in the dried state achieved by freezing followed by sublimation of water under vacuum to very low residual moisture content.

**Freezing.** The cooling of tissues to a set temperature below 0°C without the addition of a cryoprotectant.

**Frozen.** Stored in the frozen state, but without additives specifically to protect cells/matrix and/or without the controlled freezing conditions required for cryopreservation.

**Graft.** Tissues prepared for use in transplantation.

**HBc.** An abbreviation for hepatitis B core.

**HBsAg.** An abbreviation for hepatitis B virus surface antigen.

**HCV.** An abbreviation for hepatitis C virus.

**HIV.** An abbreviation for human immunodeficiency virus.

**HTLV.** An abbreviation for human T-cell lymphotropic virus.

**Hypothermic Storage.** Stored in a solution at 2 to 8°C

**Intermediate Term Preservation.** Cornea or corneal section preserved in a solution that maintains cellular and/or ultrastructure viability for 14 days. Intermediate term preservation is currently utilized at 2-8°C storage temperatures. Some types of intermediate term storage solutions are: Cornisol, Eusol-C, Kerasave, Life4°C, and Optisol GS.

**Laser Assisted Keratoplasty.** Corneal surgeries in which wound architecture is processed by laser.

a. Laser Assisted Anterior Keratoplasty. Anterior keratoplasty in which the lamellar and/or side dissection of the donor tissue is processed using a laser.

b. Laser Assisted Endothelial Keratoplasty. Endothelial keratoplasty (surgery) in which the lamellar and/or side dissection of the donor tissue is processed using a laser.

c. Laser Assisted Penetrating Keratoplasty. Penetrating keratoplasty in which the donor and recipient wound architecture are processed by laser.

**Laser Shaped.** Full-thickness cornea shaped to a specific edge profile using laser technology.

**Limbal Tissue.** Tissue bridging the junction between the cornea and sclera.

**Long Term Preservation.** Cornea or corneal section stored in a solution that is designed to maintain tissue ultrastructure for greater than 14 days and up to five years depending on the technique. Viability is not maintained. Examples are ethanol and glycerin preservation. Other media, such as albumin, may be used in conjunction with ionizing radiation to preserve the tissue ultrastructure.

**Medical Director.** See GAEBA Medical Standard B1.200.

**Microorganism.** A microscopic organism; viruses, while sometimes included in this classification, are not included here.

**Moist Chamber.** Whole eye stored at 2 to 8°C in a humid environment.

**Next of Kin.** The person(s) most closely related to a deceased individual as designated by applicable law

**ODO.** An abbreviation for organ donation organization.

**OPO.** An abbreviation for organ procurement organization.

**Organ Culture.** Stored in a nutrient medium at 28° to 37° Celsius.

**Pachymetry.** A measurement of thickness of a cornea or corneal segment (e.g., graft thickness of posterior layers processed for DSAEK).

**Package.** A labeled carton, receptacle, or wrapper containing one or more containers and accompanying labeling and package insert materials.

**Package Insert.** The written material accompanying tissue bearing further information about the tissue, directions for use, and any applicable warnings.

**Pathogen Reduction.** Describes the method of sterilization, disinfection, or decontamination of the product.

**Penetrating Keratoplasty (PK).** Full thickness cornea transplantation.

**Physical Assessment.** A limited autopsy or recent ante-mortem or post-mortem physical examination of the donor to assess for signs of a relevant communicable disease and for signs suggestive of any risk factor for a relevant communicable disease.

**Physical Examination.** See Physical Assessment.

**Plasma Dilution.** A decrease in the concentration of the donor’s plasma proteins and circulating antigens or antibodies resulting from the transfusion of blood or blood components and/or infusion of fluids, i.e., colloid(s) and/or crystalloid(s).

**Policies and Procedures Manual.** A group of standard operating procedures (SOPs) that detail the specific policies of an eye bank and the procedures used by the staff/personnel. This includes but is not limited to, procedures to assess donor eligibility; this includes operations such as; screening, recovery, processing, evaluation, testing, quarantine, labeling, storage, distribution, tracking, disposition, and recalling tissue.

**Posterior Layer.** Endothelium on Descemet membrane with or without a supporting layer of posterior stroma.

**Pre-Cut Tissue.** Corneal tissue in which lamellar or vertical dissection has been processed for surgical use, by eye bank or other organization, prior to distribution for surgical use.

**Pre-Descemet's layer Endothelial Keratoplasty (PDEK):** Transplantation of corneal endothelium, Descemet membrane, and pre-Descemet membrane.

**Pre-Distribution Shipment.** Shipment of tissue in quarantine within an establishment or between establishments (recovering eye bank to processing eye bank) of tissue that has not been released for distribution. Tissue must be shipped in quarantine.

**Preservation.** The use of chemical agents, alterations in environmental conditions or other means to prevent or retard biological or physical deterioration of ocular tissues.

**Primary Graft Failure.** Corneal edema present from the time of keratoplasty that does not clear after eight weeks and in which there is no known operative or postoperative complication or underlying recipient condition that would explain the biologic dysfunction

**Procedure.** A series of steps, which when followed, are designed to result in a specific outcome.

**Process Controls.** A system of checks and balances incorporated into standard operating procedures involving critical operations to prevent errors.

**Process Validation Studies.** The process of demonstrating a specific process or procedure will consistently produce expected results within predetermined specifications.

**Processing.** Any activity performed on the eye tissue, other than recovery, donor screening, donor testing, storage, labeling, packaging, or distribution, such as: testing for microorganisms; preparation, sterilization, steps to inactivate or remove adventitious agents; preservation for storage; manipulation/sizing and removal from storage. Any manipulation of the ocular tissue intended for transplant that involves opening a previously sealed container after recovery.

**Processing Establishment.** The entity that performs post-recovery tissue processing.

**Processing Facility Information Code.** The facility identification code of the entity that assigned the Product Code, when it is not the same facility that assigned the Donation Identification Number (DIN).

**Procurement.** See Recover(y).

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**Proficiency.** An evaluation of laboratory methods and test results that assesses the quality of standard operating procedures, equipment, supplies, and reagents, as well as the skill of the personnel performing the testing.

**QA.** An abbreviation for quality assurance.

**QC.** An abbreviation for quality control.

**Qualification.** The method of establishing confidence that equipment, reagents, and ancillary systems are capable of consistently operating within established limits and tolerances. Process performance qualification is intended to establish confidence that the process is effective and reproducible.

**Qualified.** Deemed competent by a recognized authority.

**Quality.** The conformance of ocular tissue or a process with pre-established specifications or standards.

**Quality Assurance (QA) Program.** A program that: 1) defines the policies and environment required to meet standards of quality and safety and, 2) provides confidence that the processes and tissue consistently conform to requirements for quality. Dimensions of QA may include quality control, auditing and process control, standards for personnel, facilities, procedures, equipment, testing, and record keeping activities.

**Quality Control (QC).** Specific tests defined by the eye bank’s QA Program to be performed to monitor retrieval, processing, preservation and storage, tissue quality, and test accuracy. These may include, but are not limited to performance evaluations, inspection, testing, and controls used to determine the accuracy and reliability of the eye bank’s equipment and operational procedures, as well as the monitoring of supplies, reagents, equipment, and facilities.

**Quarantine.** The identification of ocular tissue as not currently eligible for transplantation, including ocular tissue that has not yet been characterized as being eligible for transplantation. Quarantine includes the storage of such tissue in an area clearly identified for such use, or other procedures, such as automated designation, to prevent the premature release of such ocular tissue for transplantation.

**Recall.** An action taken to locate and retrieve tissue from distribution and dispensary inventories. Removal or correction of a marketed product that the FDA/other governmental or regulatory agency considers to be in violation of the laws it administers and against which the agency would initiate legal action, e.g., seizure. Recalls may be conducted on a firm’s own initiative, by regulatory agency request, or order under statutory authority.

**Receiver.** An establishment (facility or entity) or individual that accepts shipment of distributed tissues for transplantation (e.g., physicians, dentist, institutions, and other eye banks).

**Recipient.** An individual who receives an ocular tissue transplant.

**Recover(y).** The removal, acquisition, recovery, or collection of donor tissue.

**Recovery Establishment.** Entity that recovers tissue from a donor.

**Relevant Communicable Disease.** Any communicable disease relevant to transplantation of tissue in humans as defined by local regulations,

**Relevant Medical Records.** A collection of documents including a current donor risk assessment interview, a physical assessment/physical examination of the donor, laboratory test results (in addition to results of testing for required relevant communicable disease agents), relevant donor records, existing coroner and autopsy reports, as well as information obtained from any source or records that may pertain to donor eligibility regarding high risk behaviors, and clinical signs and symptoms for any relevant communicable disease agent or disease (RCDAD), and/or treatments related to medical conditions suggestive of such risk.

**Repeat Reactive.** A blood sample that is reactive on initial testing and is still reactive in at least one of two duplicate samples when the same test is repeated using the same blood sample.

**Resolution.** Adjustment, clarification, and/or correction of practices and/or procedures that results in compliance with the SOPM and/or standards.

**Responsible Party.** A person who is authorized to perform designated functions for which he or she is trained and qualified.

**Retrieval.** See Recover(y).

**Safety.** A level of quality of tissue that indicates handling according to acceptable standards and assures substantial freedom from the potential for harmful effects to recipients. The condition of being protected from risk or injury associated with occupational exposure.

**Satellite Facility.** An establishment in a location physically separate from its main facility where any activities occur that contribute to screening, recovery, transport, processing, evaluation, testing, quarantine, labeling, storage, distribution, tracking, disposition, or recall of ocular tissue under the management or direct supervision of the same corporate entity or its employee(s).

**Sclera.** Fibrous white outer part of the eye remaining after excision of the corneoscleral disc and removal of intraocular content and extraneous surface tissue.

**Secondary Carrier.** Any transporter of tissue that is not directly employed by the eye bank. Examples include, but are not limited to, contracted couriers, volunteer drivers, and commercial carriers.

**Services to Donor Families.** A defined policy or program that implements an eye bank’s recognition of the value of donation by the consenting party. These may include written communications regarding potential uses of tissue; recovery outcome information; bereavement support; provision of a copy of the Document of Consent; and/or guidance describing how to contact the eye bank if any questions arise regarding the donation. Frequency of follow-up and program maintenance is at the discretion of the Executive Director.

**Shall.** A figure of speech used interchangeably with “must.”

**Short term preservation. Eye** tissue preservation techniques that maintain viability and/or ultrastructure for less than 5 days. Examples are whole eyes preserved in a moist chamber at 2-8°C and MK medium.

**Should.** A figure of speech used to indicate a recommendation; advisory, indicating a commonly accepted activity for which there may be effective alternatives.

**SOP.** An abbreviation for standard operating procedures.

**Source Establishment (or Facility).** The entity that releases tissue following donor eligibility determination, and is responsible for maintaining donor records and evaluating adverse reaction reports

**Standard Precautions.** Guidelines recommended by the Centers for Disease Control and Prevention for reducing the risk of transmission of bloodborne and other pathogens. Standard precautions apply to (1) blood; (2) all body fluids, secretions, and excretions except sweat, regardless of whether or not they contain blood; (3) non-intact skin; and (4) mucous membranes. Standard Precautions includes hand hygiene, and the use of appropriate personal protective equipment such as gloves, gown, mask, eye protection, or face shield, whenever touching or exposure to patients’ body fluids is anticipated.

**Sterile.** The absence of detectable, viable, microorganisms.

**Sterilization.** A validated method used to render instrumentation and ocular tissue free from viable microorganisms, including spores.

**Store or Storage.** The maintenance of ocular tissue for future use.

**Storage Establishment.** The entity that stores tissue at any time prior to distribution to the end user.

**Storage Solution.** Specifies the solution in which the tissue is stored.

**Summary of Records.** A condensed version of the required testing and screening records that contains the identity of the testing laboratory, the listing and interpretation of all required infectious disease tests, a listing of the documents reviewed as part of the relevant medical records, and the name of the person or establishment determining the eligibility of the ocular tissue for transplantation.

**Time of Death.** For purposes of eye donation, the time of death is the cessation of heartbeat, cardiac death, asystole, cross-clamp, last known alive (LKA), or it can be the time of death established by core temperature, when applicable and with appropriate documentation from a medical professional.

**Tissue.** A functional group of ocular cells, such as cornea, sclera. Tissues may be transplanted as viable cells or otherwise preserved or fixed. Tissue does not include perfusable organs for transplantation.

**Tissue Bank.** An entity that provides or engages in one or more services involving tissue from living or deceased individuals for transplantation, research, and/or educational purposes. These services include but are not limited to assessing donor suitability (including screening), recovery, processing, evaluation, testing, quarantine, labeling, storage, distribution, tracking, disposition, and recall of tissue.

**Tissue Evaluation.** Microscopic or other analysis of donor eye tissue to determine if tissue is suitable for intended use(s).

**Tissue Type.** Refers to whole eye or a specific portion of the eye, such as cornea or sclera.

**Tolerance Limits.** The limits that define a range of acceptable values established for each testing procedure that, when exceeded, require the implementation of corrective actions designed to produce results within the acceptable range in future tests.

**Traceability.** The act or ability to locate ocular tissue during any step of its recovery, processing, evaluation, testing, quarantine, labeling, storage, distribution, disposition, or recall. It includes the capacity to identify the surgeon, consignee, or medical facility receiving the tissue and the ability of the surgeon, consignee, or medical facility to identify the storage, recipient, or final disposition of the tissue.

**Tracking.** The act or ability to locate individual tissue during any step of its recovery, processing, evaluation, testing, quarantine, labeling, storage, distribution, disposition, and recalling. It includes the capacity of the distributing eye bank to identify the consignee and the consignee to identify the recipient.

**Transplantation.** The transfer of tissue to a recipient.

**Transplant Program.** An organization of medical personnel and allied health care professionals, operating in one or more transplant centers, with the responsibility for the transplantation of one of more types of tissues, and/or organs.

**Transport Medium.** Any microbiological medium capable of maintaining cellular viability during the transport of ocular tissue.

**Universal Precautions.** An approach to infection control, in which all human blood, certain body fluids, as well as fresh tissues and cells of human origin are handled as if they are known to be infected with HIV, HBV, and/or other blood-borne pathogens.

**Validation.** The method of establishing documented evidence that provides a high degree of assurance that specific process will consistently produce the predetermined outcome.

**Variance.** A departure from Standards that is pre-approved by the Medical Director, Responsible Party, or governing authority prior to implementation.

**Verification.** The confirmation by examination and provision of objective evidence that specified requirements have been fulfilled.

**Wet Ice Temperatures.** Temperatures ranging from 1-10°C.

**Withdrawal (or Market Withdrawal).** Removal or correction of a distributed product by an eye bank that involves a minor violation that would not be subject to legal action by governmental or regulatory agency, or that involves no violation. Does not involve a relevant communicable disease.

**Appendix I-Communicable Disease**

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Medical Standards Appendix I: Relevant Communicable Disease Agents and Diseases

Introduction

As referenced in D1.120, this appendix contains definition of relevant communicable disease and disease agents. The guidance does not establish legally enforceable responsibilities, but describes the current thinking on the contained topics and should be viewed only as recommendations unless specific regulatory or statutory requirements are cited. The use of the word “should” means that something is suggested or recommended, but not required.

Guidance

I. Relevant communicable disease and disease agents

a. The following communicable diseases and disease agents are relevant for ocular tissue:

i. Human immunodeficiency virus (HIV), types 1 and 2;

ii. Hepatitis B virus (HBV);

iii. Hepatitis C virus (HCV);

iv. Human transmissible spongiform encephalopathy (TSE); including Creutzfeldt-Jakob disease (CJD); and

v. Treponema pallidum (syphilis).

II. A communicable disease agent is relevant if it is one

a. For which there may be a risk of transmission by ocular tissue, either to the recipient of the ocular tissue or to those people who may handle or otherwise come in contact with the ocular tissue, such as medical personnel, because the disease agent or disease:

i. is potentially transmissible by ocular tissue; and

ii. either (1) has sufficient incidence and/or prevalence to affect the potential donor population, or (2) may have

been released accidentally or intentionally in a manner that could place potential donors at risk of infection;

b. That could be fatal or life-threatening, could result in permanent impairment of a body function or permanent damage to body structure, or could necessitate medical or surgical intervention to preclude permanent impairment of body function or permanent damage to a body structure;and

c. For which appropriate screening measures have been developed and/or an appropriate screening test for donor specimens has been licensed, approved, or cleared for such use and is available.

d. Examples of RCDADs considered as relevant include, but are not limited to:

i. West Nile Virus

ii. Sepsis

iii. Vaccinia

**Appendix II-Contraindications Guidance**

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I. Risk Factors

Following is a list of conditions and behaviors that increase the donor’s relevant communicable disease risk. Except as noted in this section, you should determine to be ineligible any potential donor who exhibits one or more of the following conditions or behaviors.

a. Men who have had sex with another man in the preceding 5 years (risk factor for HIV and Hepatitis B).

b. Persons who have injected drugs for a non-medical reason in the preceding 5 years, including intravenous, intramuscular, or subcutaneous injections (risk factor HIV, Hepatitis B and Hepatitis C).

c. Persons who have engaged in sex in exchange for money or drugs in the preceding 5 years (risk factor for HIV, Hepatitis B and Hepatitis C).

d. Persons who have had sex in the preceding 12 months with any person described (as above in I.a-c) or with any person who has HIV infection, including a positive or reactive test for HIV virus, hepatitis B infection, or clinically active (symptomatic) hepatitis C infection.

e. Persons who have been exposed in the preceding 12 months to known or suspected HIV, HBV, and/or HCV-infected blood through percutaneous inoculation (e.g., needle stick) or through contact with an open wound, non- intact skin, or mucous membrane.

f. Children born to mothers with or at risk for HIV infection, if the child is 18 months of age or younger, or if the child was breast-fed within the preceding 12 months.

g. Persons who have been in juvenile detention, lock up, jail or prison for more than 72 consecutive hours in the preceding 12 months (risk factor for HIV, Hepatitis B and Hepatitis C).

h. Persons who have lived with (resided in the same dwelling) another person who has hepatitis B or clinically active (symptomatic) hepatitis C infection in the preceding 12 months.

i. Persons who have undergone tattooing, ear piercing or body piercing in the preceding 12 months, in which sterile procedures were not used, e.g., contaminated instruments and/or ink were used, or shared instruments that had not been sterilized between uses were used.

j. Persons who have had a past diagnosis of clinical, symptomatic viral hepatitis after their 11th birthday, unless evidence from the time of illness documents that the hepatitis was identified as being caused by hepatitis A virus, Epstein-Barr Virus (EBV), or cytomegalovirus (CMV).

k. Persons who are deceased and have a documented medical diagnosis of sepsis or have documented clinical evidence consistent with a diagnosis of sepsis that is not explained by other clinical conditions at the time of death. An eye bank should make a determination on how to routinely handle situations of clinical history proximal to death in which sepsis was suspected at the time of admission or part of a differential diagnosis during admission in which the patient may have been shown through clinical data not to be septic prior to death.

l. Persons who have had smallpox vaccination (vaccinia virus) in the preceding 8 weeks should be evaluated as follows:

For persons who had no vaccinia complications (including eczema vaccinatum, generalized vaccinia, progressive vaccinia, postvaccinial encephalitis, or vaccinial keratitis):

1. You should defer the donor until after the vaccination scab has separated spontaneously, or for 21 days post-vaccination, whichever is the later date, and until the physical examination or physical assessment includes a confirmation that there is no scab at the vaccination site.

**Appendix II-Contraindications Guidance**

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2. In cases where a scab was removed before separating spontaneously, you should defer the donor for two months after vaccination.

3. In cases where the eye bank cannot determine how the scab separated, you should defer if the vaccination was less than 21 days ago. Deferral is not necessary if the scab is not visible upon external examination and the vaccination was greater than 21 days ago.

4. For persons who have experienced vaccinia complications, you should defer the donor until 14 days after all vaccinia complications have completely resolved. Deferral is not necessary if the date of vaccinia complication resolution is indeterminate and there are no visible signs of vaccinia complications.

5. Persons who acquired a clinically recognizable vaccinia virus infection by contact with someone who received the smallpox vaccine (i.e., touching the vaccination area or the scab, including the covering bandages, or touching clothing, towels, or bedding that might have come into contact with an unbandaged vaccination area or scab).

6. For cadaveric donors who have received a smallpox vaccination or had close contact with the smallpox vaccination site of someone else, you should examine the skin and defer if a scab or other signs of vaccinia are present.

7. You should defer persons who developed other complications of vaccinia infection acquired through contact with a vaccine recipient until 14 days after all vaccinia complications have completely resolved. Deferral is not necessary if the date of vaccinia complication resolution is indeterminate and there are no visible signs of vaccinia complications. Deferral is not necessary for anyone who did not develop vaccinia complications.

m. Persons who have had a medical diagnosis or suspicion of WNV infection (based on symptoms and/or laboratory results or confirmed WNV viremia) you should defer for 120 days following diagnosis or onset of illness, whichever is later.

n. Persons who have tested positive or reactive for WNV infection in the preceding 120 days.

o. Persons who have been treated for or had syphilis within the preceding 12 months. We do not recommend deferral of donors if evidence is presented that the treatment occurred more than 12 months ago and was successful.

p. Persons who have been diagnosed with vCJD or any other form of CJD.

q. Persons who have been diagnosed with dementia or any degenerative or demyelinating disease of the central nervous system or other neurological disease of unknown etiology. Examples include Parkinson, amyotrophic lateral sclerosis, multiple sclerosis, Alzheimer disease, Guillain-Barre, and Chronic Inflammatory Demyelinating Polyneuropathy (CIPD). Potential donors who have a diagnosis of delirium (e.g., delirium caused by toxic/metabolic diseases or recent head trauma) would not necessarily be

considered to have a diagnosis of dementia and should be evaluated by the Medical Director. (Ocular tissue from donors with dementia confirmed by gross and microscopic examination of the brain to be caused by cerebrovascular accident or brain tumor and who are confirmed not to have evidence of TSE on microscopic examination of the brain may be acceptable based on an evaluation by the Medical Director).

r. Persons who are at increased risk for CJD. Donors are considered to have an increased risk for CJD if they have received a non-synthetic dura mater transplant, human pituitary-derived growth hormone, or have one or more blood relatives diagnosed with CJD.

s. Persons who have a history of CJD in a blood relative unless the diagnosis of CJD was subsequently found to be an incorrect diagnosis, the CJD was iatrogenic, or the laboratory testing (gene sequencing) shows that the donor does not have a mutation associated with familial CJD.

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t. If the eye bank is not testing for HIV I/II using a test kit specifically labeled as sensitive for detection of HIV group O antibodies, then deferral includes persons or their sexual partners who were born or lived in certain countries in Africa (Cameroon, Central African Republic, Chad, Congo, Equatorial Guinea, Gabon, Niger, or Nigeria) after 1977 (risk factor for HIV group O).

u. If the eye bank is not testing for HIV I/II using a test kit specifically labeled as sensitive for detection of HIV group O antibodies, then deferral includes persons who have received a blood transfusion or any medical treatment that involved blood in Cameroon, Central African Republic, Chad, Congo, Equatorial Guinea, Gabon, Niger, or Nigeria, after 1977 (risk factor for HIV group O).

v. Persons who are xenotransplantation product recipients or intimate contacts of a xenotransplantation product recipient. Xenotransplantation is any procedure that involves the transplantation, implantation, or infusion into a human recipient of either: (1) live cells, tissues, or organs from a nonhuman animal source; or (2) human body fluids, cells, tissues, or organs that have had ex vivo contact with live nonhuman animal cells, tissues, or organs. Xenotransplantation products include live cells, tissues, or organs used in xenotransplantation. Biological products, drugs, or medical devices sourced from nonliving cells, tissues or organs from nonhuman animals, including but not limited to porcine insulin and porcine heart valves, are not considered xenotransplantation products. Xenotransplantation product recipient means a person who undergoes xenotransplantation. Intimate contact of a xenotransplantation product recipient means a person who has engaged in activities that could result in intimate exchange of body fluids, including blood or saliva, with a xenotransplantation product recipient. Intimate partners of recipients of the xenotransplantation product Epicel™ need not be deferred.

II. Clinical Evidence

Except as noted in this section), you should determine to be ineligible any potential donor who exhibits one or more of the following examples of clinical evidence of relevant communicable disease.

a. HIV Infection

1. A prior positive or reactive screening test for HIV;

2. Unexplained weight loss;

3. Unexplained night sweats;

4. Blue or purple spots on or under the skin or mucous membranes typical of Kaposi's sarcoma;

5. Disseminated lymphadenopathy (swollen lymph nodes) for longer than one month;

6. Unexplained temperature of >100.5oF (38.6oC) for more than 10 days;

7. Unexplained persistent cough or shortness of breath;

8. Opportunistic infections;

9. Unexplained persistent diarrhea; and/or

10. Unexplained persistent white spots or unusual blemishes in the mouth.

b. Hepatitis Infection:

1. A prior positive or reactive screening test for hepatitis B virus or hepatitis C virus;

2. Unexplained jaundice;

**Appendix II-Contraindications Guidance**

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3. Unexplained hepatomegaly; and/or

4. Past diagnosis of clinical, symptomatic viral hepatitis after the 11th birthday, unless evidence from the time of illness documents that the hepatitis was identified as caused by hepatitis A virus, EBV, or CMV.

Note: Records of the following laboratory data might assist you in making the donor-eligibility determination in the face of an inconclusive history of hepatitis infection: alanine aminotransferase (ALT), aspartate aminotransferase (AST), bilirubin or prothrombin time. If these tests are abnormal, but a cause other than viral hepatitis was established, we do not recommend that you defer the donor.

c. Syphilis Infection:

1. Persons who have had or have been treated for syphilis, in the preceding 12 months. We do not recommend deferral of donors who have had or have been treated for syphilis more than 12 months ago, if evidence is presented that treatment occurred more than 12 months ago and was successful.

d. Vaccinia Infection:

1. Recent smallpox vaccination;

2. Eczema vaccinatum;

3. Vesicular rash indicative of generalized vaccinia in a person who has had recent smallpox immunization or who is a contact of someone with recent smallpox immunization.

4. Progressive necrosis in an area of vaccination consistent with vaccinia necrosum;

5. Postvaccinial encephalitis; and/or

6. Vaccinial keratitis.

e. WNV Infection:

1. Mild symptoms might include fever, headache, body aches, or eye pain;

i. mild symptoms might also occasionally be accompanied by a skin rash on the trunk of the body; or

ii. swollen lymph glands.

2. Severe illness;

i. severe illness might include encephalitis, meningitis, meningoencephalitis, and acute flaccid paralysis;

ii. signs and symptoms of severe illness might include headache, high fever, neck stiffness, stupor, disorientation, coma, tremors, convulsions, and muscle weakness or paralysis.

f. Sepsis (including, but not limited to, bacteremia, septicemia, sepsis syndrome, systemic infection, systemic inflammatory response syndrome (SIRS), or septic shock):

1. Clinical evidence of infection; and

2. Two or more of the following systemic responses to infection if unexplained:

i. Temperature of >100.4o F (38o C);

ii. Heart rate >90 beats/min;

**Appendix II-Contraindications Guidance**

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iii. Respiratory rate >20 breaths/min or PaCO2 <32; or

iv. WBC >12,000 cells/mm3, <4,000 cells/mm3, or >10% immature (band) forms.

3. More severe signs of sepsis include unexplained hypoxemia, elevated lactate, oliguria, altered mentation, and hypotension.

4. Positive (pre-mortem) blood cultures might be associated with the above signs.

III. Physical Evidence

You should determine ineligible for transplant any potential donor who exhibits one or more of the following examples of physical evidence of RCDADs or high-risk behavior associated with these RCDADs.

a. Physical evidence for risk of sexually transmitted diseases such as genital ulcerative disease, herpes simplex, chancroid (you should consider these signs in light of other information obtained about the donor in making a donor eligibility determination) (seen in HIV, Hepatitis B virus, Chlamydia trachomatis, and Neisseria gonorrhea).

b. Physical evidence for risk of, or evidence of syphilis.

c. For a male donor, physical evidence of anal intercourse including perianal condyloma (seen in HIV and Hepatitis B).

d. Physical evidence of nonmedical percutaneous drug use such as needle tracks; your examination should include examination of tattoos, which might be covering needle tracks (seen in HIV, Hepatitis B and Hepatitis C).

e. Physical evidence of recent tattooing, ear piercing, or body piercing. Persons who have undergone tattooing, ear piercing, or body piercing in the preceding 12 months, in which sterile procedures were not used (e.g., contaminated instruments and or/ink were used), or instruments that had not been sterilized between uses were used (seen in HIV, Hepatitis B and Hepatitis C).

f. Disseminated lymphadenopathy (seen in HIV).

g. Unexplained oral thrush (seen in HIV).

h. Blue or purple spots consistent with Kaposi's sarcoma (seen in HIV).

i. Unexplained jaundice, hepatomegaly, or icterus (seen in Hepatitis B and Hepatitis C). Note: Hepatomegaly may not be apparent in a physical assessment unless an autopsy is performed.

j. Physical evidence of sepsis, such as unexplained generalized rash or fever.

k. Large scab consistent with recent history of smallpox immunization.

l. Eczema vaccinatum (seen in vaccinia).

m. Generalized vesicular rash (generalized vaccinia).

n. Severely necrotic lesion consistent with vaccinia necrosum.

o. Corneal scarring consistent with vaccinial keratitis.

**Appendix III- Donor Eligibility Determination Guidance**

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I. The Process of Donor Eligibility Determinations:

a. A donor-eligibility determination is a conclusion by a responsible person (Medical Director or Designee) that an ocular tisue donor is either eligible or ineligible to donate for clinical use, based on the results of donor screening and testing A donor is eligible for clinical use only if screening shows that the donor is free from risk factors for, and clinical evidence of, infection due to relevant communicable disease agents and diseases (RCDADs), and is free from communicable disease risks associated with xenotransplantation; and test results for relevant communicable disease agents are negative or nonreactive, except for non-treponemal screening tests for syphilis.

b. You must establish and maintain procedures for all steps that you perform in testing, screening, determining donor eligibility, and complying with all other requirements of relevant regulatory agencies. A responsible person must review and approve all procedures before their

Implementation. These procedures must be readily available to personnel in the area where the procedures are performed, or if this is not practical, in a nearby area. You may use appropriate standard procedures developed by another organization, provided that you have verified that the procedures are consistent with and at least as stringent as the relevant regulatory requirements.

c. At the time a departure occurs, you must record and justify that departure from a procedure relevant to preventing risks of communicable disease transmission. Before making available for distribution ocular tissue from a donor whose eligibility is to be determined under such a departure, a responsible person must determine that the departure did not increase the risk of communicable disease transmission. A departure is a change from an established procedure, including Standard Operating Procedure (SOP), which occurs before the ocular tissue is distributed, and is consistent with applicable regulations and standards.

A deviation is an event that inconsistent with applicable regulations, standards, or established specifications, and is discovered after tissue has been released for surgical use and is either unexpected or unforeseeable.

II. Records to Accompany Ocular Tissue After Determination of Donor Eligibility

a. A distinct identification code (such as an alphanumeric code) affixed to the ocular tissue container, that relates the ocular tissue to the donor and to all records pertaining to the ocular tissue and does not include an individual’s name, social security number, or medical record number

b. A statement that based on the results of screening and testing, the donor is determined to be eligible

c. A summary of the records used to make the donor-eligibility determination,including:

i. A statement that the communicable disease testing was performed by a laboratory or laboratories certified to perform such testing on human specimens by any relevant regulatory agencies.

ii. A listing and interpretation of the results of all tests performed for relevant communicable disease agents or diseases

iii. The name and address of the establishment that made the donor- eligibility determination

III. Information Sources for Review for Donor Eligibility Determination

a. When you screen an ocular donor, you must review relevant medical records for risk factors for, and clinical evidence of RCDADS (see Appendix II). Relevant medical records means a collection of documents that includes: (1) a current donor medical history interview; (2) a current report of the physical assessment of a cadaveric donor or the physical examination of a living donor; and (3) other available records.

**Appendix III- Donor Eligibility Determination Guidance**

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i. The donor medical history interview is a documented dialogue concerning the donor's medical history and relevant social behavior with a living donor or if the donor is not living or is unable to participate in the interview, then with one or more individuals who can provide the information sought. These individuals might be the donor's next of kin, the nearest available relative, a member of the donor's household, an individual with an affinity relationship with the donor (e.g., caretaker, friend, partner), or the donor's primary treating physician.

ii. The purpose of the physical assessment of a cadaveric donor or the physical examination of a living donor is to assess for physical signs of a relevant communicable disease and for signs suggestive of any risk factor for such a disease (reference Appendix II). For a cadaveric donor, the physical assessment means a limited autopsy, or a recent antemortem or postmortem physical examination. For living donors, you may examine only those parts of the body that are necessary to evaluate for RCDADs based upon relevant donor history that has been obtained during the interview and review of available records. You may rely on records of a recent report of a physical examination by other health care professionals. Because this is a step in determining donor eligibility, you must establish and maintain standard operating procedures (SOPs) for the conduct of the physical assessment or physical examination.

iii. If they are available, other records that also meet the definition of relevant medical records include

• Laboratory test results (other than the results of testing required for the donor-eligibility determination);

• Medical records;

• Coroner and autopsy reports; and

• Records or other information received from any source pertaining to risk factors for relevant communicable disease

(e.g., social behavior, clinical signs and symptoms of relevant communicable disease, and treatments related to medical conditions suggestive of risk for relevant communicable disease). Examples of these records include medical examiner reports, police records, and information from other tissue or medical establishments, if applicable.

NOTE: In the absence of autopsy reports available in time for release of transplant use of corneas, an eye bank should review the presumed cause of death and other relevant preliminary autopsy findings, then review the autopsy report when it becomes available after release of tissue. If new information in the final report indicates that the donor is ineligible, you should consider notifying the consignees of the distributed tissue and submit a report to any relevant regulatory agency as dictated by local requirements.

**Appendix IV-Testing**

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I. Donor Testing

a. You must test all donors of ocular tissue for the RCDADs listed belowYou must use an approved or cleared screening test in accordance with the manufacturer’s instructions. You must use a donor screening test specifically labeled for cadaveric specimens instead of a more generally labeled donor screening test when applicable and when available. Testing must be performed at a testing facility that has been certified under the requirements the relevant regulatory agency. You must determine a donor ineligible whose specimen tests reactive for a screening test for a relevant communicable disease agent, except for the Treponema pallidum exception below.

Testing is required for the following RCDADs:

i. Human immunodeficiency virus, type 1 (HIV-1)

ii. Human immunodeficiency virus, type 2 (HIV-2)

iii. Hepatitis B virus (HBV)

iv. Hepatitis C virus (HCV)

v. Treponema pallidum (Syphilis)

b. Recommended screening tests

i. HIV, type 1 (licensed screening test either for anti-HIV-1 or combination test for anti-HIV-1 and anti-HIV-2; and licensed screening NAT test for HIV-1, or combination NAT); (if the screening test used is not labeled as sensitive for group O antibodies, donors must be evaluated for risk factors associated with HIV group O, described in Appendix II, I.x-y.)

ii. HIV, type 2 (licensed screening test either for anti-HIV-2 or combination test for anti-HIV-1 and anti-HIV-2)

iii. HBV (licensed screening test for Hepatitis B surface antigen (HBsAg); for total antibody to Hepatitis B core antigen (anti-HBc) (IgG and IgM); and licensed NAT donor screening test for HBV.

iv. HCV (FDA-licensed screening test for anti-HCV; and FDA-licensed screening NAT test for HCV, or combination NAT)

v. Treponema pallidum (licensed, approved, or cleared screening test for syphilis. As an exception for syphilis test results,you may determine to be eligible a donor whose specimen tests positive or reactive on a non-treponemal screening test for syphilis and negative or nonreactive on a specific treponemal confirmatory test, so long as all other required testing and screening are negative or nonreactive. A donor whose specimen tests positive or reactive on either a specific treponemal confirmatory test for syphilis or on a treponemal screening test is not eligible. If a cadaveric specimen is too hemolyzed to interpret the Rapid Plasma Reagin (RPR) test result, you should use another treponemal screening test.

II. Assessment of Donor Specimens for Testing Suitability

You must collect the donor specimen for testing at the same time as tissue is recovered from the donor, or within seven days before or after the recovery of tissue. You may use a premortem specimen to test a cadaveric donor, as long as the specimen is collected within that timeframe. Although there is no requirement that

specifies when to test the collected specimen, you should perform testing as soon as possible after collection and in accordance with the time limits stated in the manufacturer’s instructions for use of the test kit.

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For adult donors who have suffered blood loss sufficient to require fluid replacement, certain volumes of transfusions and/or infusions should be suspected of affecting test results. Blood loss might occur internally or externally. For donors 12 years of age or younger, you should suspect that any transfusion or infusion might affect

test results regardless of blood loss. There might be other clinical situations involving transfusion or infusion that should also be suspected of affecting test results. Autologous blood removed pre-operatively or peri-operatively and reinfused during the same surgical procedure would not need to be included in plasma dilution calculations.

a. For adult donors you must suspect plasma dilution sufficient to affect the results of communicable disease agent testing where blood loss is known or suspected in a donor over 12 years of age in any of the following situations:

1. The donor received a transfusion or infusion of more than 2000 milliliters of blood or colloids either:

i. Within the 48 hours immediately preceding the collection of a pre-mortem specimen for testing; or

ii. Within the 48 hours immediately preceding death, if the specimen for testing is collected post-mortem, whichever occurred earlier.

2. The donor received more than 2000 milliliters of crystalloids within either:

i. The one hour immediately preceding the collection of a pre-mortem specimen for testing; or

ii. Within the one hour immediately preceding death, if the specimen for testing is collected post-mortem, whichever occurred earlier.

3. The donor received more than 2000 milliliters of any combination of whole blood, red blood cells, colloids, and/or crystalloids within the applicable time frames set out in (1) and (2) in this section.

b. For pediatric donors you must suspect plasma dilution sufficient to affect the results of communicable disease agent testing, regardless of the presence or absence of blood loss, in a donor 12 years of age or under, in any of the following situations.

1. Any transfusion of blood or colloids:

i. Within the 48 hours immediately preceding the collection of a pre-mortem specimen for testing; or

ii. Within the 48 hours immediately preceding death, if the specimen is collected post-mortem, whichever occurred earlier.

2. Any crystalloids:

i. Within the one hour immediately preceding the collection of a pre-mortem specimen for testing; or

ii. Within the one hour immediately preceding death, if the specimen is collected post-mortem, whichever occurred earlier.

c. There may be other clinical situations that may cause reasonable suspicion of

a specimen to be diluted. As the establishment that collects donor specimens for testing, you might be aware of additional circumstances in which plasma dilution might affect test results. Your SOPs should identify any additional circumstances where you believe plasma dilution might have occurred, and you should use a pre-transfusion/infusion specimen or apply an algorithm in those instances.

1. Example 1: A donor who has previously had blood loss, stabilizes, then expires, but has received fluids in the 48 hours before specimen collection.

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2. Example 2: A donor who is obese.

3. Example 3: A donor who in the absence of bleeding may have received large amounts of infusions which the medical director or designee believes may affect test results.

4. Example 4: A donor who weighs less than 45 kilograms or more than 100 kilograms.

d. As part of establishing procedures for all steps in testing establishments making donor eligibility determinations must have SOPs that define those elements necessary to determine whether a pre-transfusion/infusion blood specimen is adequate for infectious disease testing (e.g., the amount of hemolysis, storage conditions, and age of the specimen). Testing laboratories must perform tests in accordance with the manufacturer’s instructions,including any instructions concerning factors that might affect specimen stability.

e. An appropriate algorithm must evaluate the fluid volumes administered in the 48 hours before collecting the specimen from the donor and show that plasma dilution sufficient to affect test results has not occurred

Definitions:

Blood component means a product containing a part of human blood separated by physical or mechanical means

Colloid means: (1) a protein or polysaccharide solution, such as albumin, dextran, or hetastarch, that can be used to increase or maintain osmotic (oncotic) pressure in the intravascular compartment; or (2) blood components such as plasma and platelets

Crystalloid means an isotonic salt and/or glucose solution used for electrolyte replacement or to increase intravascular volume, such as saline solution, Ringer's lactate solution, 5 percent dextrose in water, or total parenteral nutrition (TPN).

Plasma dilution means a decrease in the concentration of the donor's plasma proteins and circulating antigens or antibodies resulting from the transfusion of blood or blood components and/or infusion of fluids.