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Subject: Bio-Engineered Skin and Soft Tissue Substitutes; Amniotic Membrane and Amniotic Fluid

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DESCRIPTION:

Bio-engineered skin and soft tissue substitutes may be derived from human tissue (autologous or allogeneic), non-human tissue (xenographic), synthetic materials, or a composite of these materials. Bioengineered skin and soft tissue substitutes are being evaluated for a variety of conditions, including breast reconstruction and healing lower-extremity ulcers and severe burns. Acellular dermal matrix (ADM) products are also being evaluated for soft tissue repair.

Bioengineered skin and soft tissue substitutes may be either acellular or cellular. Acellular products (eg, dermis with cellular material removed) contain a matrix or scaffold composed of materials such as collagen, hyaluronic acid, and fibronectin. Acellular dermal matrix (ADM) products can differ in a number of ways, including as species source (human, bovine, porcine), tissue source (eg dermis, pericardium, intestinal mucosa), additives (eg antibiotics, surfactants), hydration (wet, freeze-dried), and required preparation (multiple rinses, rehydration). Cellular products contain living cells such as fibroblasts and keratinocytes within a matrix. The cells contained within the matrix may be autologous, allogeneic, or derived from other species (eg, bovine, porcine). Skin substitutes may also be composed of dermal cells, epidermal cells, or a combination of dermal and epidermal cells, and may provide growth factors to stimulate healing. Bioengineered skin substitutes can be used as either temporary or permanent wound coverings.

There are a large number of potential applications for artificial skin and soft tissue products. One large category is nonhealing wounds, which potentially encompasses diabetic neuropathic ulcers, vascular insufficiency ulcers, and pressure ulcers. A substantial minority of such wounds do not heal adequately with standard wound care, leading to prolonged morbidity and increased risk of mortality. For example, nonhealing lower-extremity wounds represent an ongoing risk for infection, sepsis, limb amputation, and

death. Bioengineered skin and soft tissue substitutes have the potential to improve rates of healing and reduce secondary complications.

Other situations in which bioengineered skin products might substitute for living skin grafts include certain postsurgical states (eg, breast reconstruction) in which skin coverage is inadequate for the procedure performed, or for surgical wounds in patients with compromised ability to heal. Second- and third-degree burns are another indication in which artificial skin products may substitute for auto- or allografts. Certain primary dermatologic conditions that involve large areas of skin breakdown (eg, bullous diseases) may also be conditions in which artificial skin products can be considered as substitutes for skin grafts. ADM products are also being evaluated in the repair of other soft tissues including rotator cuff repair, following oral and facial surgery, hernias, and other conditions.

Amniotic Membrane and Amniotic Fluid

Several commercially available forms of human amniotic membrane (HAM) and amniotic fluid can be administered by patches, topical application, or injection. Amniotic membrane and amniotic fluid are being evaluated for the treatment of various conditions, including chronic full thickness diabetic lower extremity ulcers, venous ulcers, knee osteoarthritis, plantar fasciitis, and ophthalmic conditions.

POSITION STATEMENT:

Note: This guideline does not address the use of meshes or patches of non-biologic origin used for standard repair procedures such as hernia repairs.

Bio-Engineered Skin and Soft Tissue Substitutes

Breast reconstructive surgery using allogeneic acellular dermal matrix products* (i.e. AlloDerm[®], AlloMend[®], Cortiva[®] (AlloMax[™]), DermACELL[™], DermaMatrix[™], FlexHD[®], FlexHD[®] Pliable[™], GraftJacket[®]) **meets the definition of medical necessity** for **ONE** of the following indications:

- when there is insufficient tissue expander or implant coverage by the pectoralis major muscle and additional coverage is required, **OR**
- when there is viable but compromised or thin post-mastectomy skin flaps that are at risk of dehiscence or necrosis, **OR**
- the inframammary fold and lateral mammary folds have been undermined during mastectomy and re-establishment of these landmarks is needed.

Treatment of chronic, non-infected, full-thickness diabetic lower extremity ulcers using the following tissue-engineered skin substitutes **meets the definition of medical necessity:**

- AlloPatch^{®*}
- Apligraf^{®**}
- Dermagraft^{®**}
- Integra[®] Omnigraft Dermal Regeneration Matrix (also known as Omnigraft)
- Integra[™] Flowable Wound Matrix
- Primatrix[®]
- Primatrix[®] Dermal Repair Scaffold.

Treatment of chronic, noninfected, partial- or full-thickness lower extremity skin ulcers due to venous insufficiency, which have not adequately responded following a one-month period of conventional ulcer therapy, using the following tissue-engineered skin substitutes **meets the definition of medical necessity**:

- Apligraf^{®**}
- Oasis[™] Wound Matrix^{***}.

Treatment of dystrophic epidermolysis bullosa using the following tissue-engineered skin substitutes **meets the definition of medical necessity**:

- OrCel[™] (for the treatment of mitten-hand deformity when standard wound therapy has failed and when provided in accordance with the Humanitarian Device Exemption (HDE) specifications of the FDA)^{****}.

Treatment of second- and third-degree burns using the following tissue-engineered skin substitutes **meets the definition of medical necessity**:

- Epicel[®] (for the treatment of deep dermal or full-thickness burns comprising a total body surface area of greater than or equal to 30% when provided in accordance with the HDE specifications of the FDA)^{****}
- Integra[®] Dermal Regeneration Template^{™**}
- Primatrix[®] Dermal Repair Scaffold.

* Banked Human Tissue.

** FDA premarket approval.

*** FDA 510(k) clearance.

**** FDA-approved under an HDE.

All other uses of the bio-engineered skin and soft tissue substitutes listed above **do not meet the definition of medical necessity**.

Amniotic Membrane and Amniotic Fluid

Treatment of nonhealing diabetic lower-extremity ulcers using the following human amniotic membrane products **meets the definition of medical necessity**:

- Affinity[®]
- AmnioBand[®] Membrane
- Biovance[®]
- Epicord[®]
- Epifix[®]
- Grafix[™].

All other indications not listed above are considered **experimental or investigational**, including but not limited to treatment of lower extremity ulcers due to venous insufficiency. The evidence is insufficient to determine the effects of the technology on health outcomes.

The following is considered **experimental or investigational** for all indications, including but not limited to treatment of osteoarthritis and plantar fasciitis:

- Injection of micronized or particulated human amniotic membrane
- Injection of human amniotic fluid.

The evidence is insufficient to determine the effects of the technology on health outcomes.

All other bio-engineered skin substitutes, soft tissue substitutes, amniotic membranes and amniotic fluids are considered **experimental or Investigational**, including, but not limited to:

- ACell® UBM Hydrated/Lyophilized Wound Dressing
- ActiveBarrier® 45, 200, 2000
- Allogen
- AlloSkin™
- AlloSkin™ AC
- AlloSkin™ RT
- Allowrap™
- Alphaplex™ with MariGen Omega3™
- Altrazeal™ Transforming Powder Dressing
- AmnioAMP-MP
- Amnioarmor™
- AmnioBand™ Particulate
- Amnion bio or Axomembrane
- AmnioClear™
- Amniocore™
- Amniocyte
- AmnioExCel™
- Amnio-maxx or Manio-maxx lite
- AmnioFix®
- AmnioFix® Injectable
- AmnioGen™ 45
- AmnioGen™ 200
- AmnioGen™ A
- AmnioGen™ C

- AmnioGraft®
- AmnioMatrix®
- AmnioPro® 45
- AmnioPro® 200
- AmnioPro® Flow
- Amniorepair or AltiPly
- Amniotext and Amniotext Patch
- AmnioVisc™
- Amniowound
- AmnioWrap2™
- Amniply
- Aongen™ Collagen Matrix
- Architect® ECM, PX, FX
- Artacent® Cord, Wound
- ArthroFlex™ (FlexGraft)
- Articent ac flowable, patch
- Ascent
- Atlas Wound Matrix
- Avagen Wound Dressing
- Avaulta Plus™
- AxoGuard® Nerve Protector (AxoGen)
- Axolotl ambient or Axolotl Cryo
- Biobrane®/Biobrane-L
- Bio-ConneKt® Wound Matrix
- BioDDryFlex®
- BioDfence™
- BioDfence Dryflex™/BioDfactor™
- BioDMatrix™
- BioNextPATCH
- BioRenew® 45
- BioRenew® 200
- BioRenew® Flow
- BioSkin® 45
- BioSkin® 200
- BioSkin® Flow

- BioWound, BioWound Plus™, BioWound XPlus™
- carePATCH
- CellerateRX® (CRXa)
- Cellesta/Cellesta duo, Cord, flowable
- Clarix®
- Clarix® Flo
- Cogenex amniotic membrane
- CollaCare®
- CollaCare® Dental
- Collagen Sponge
- Collagen Wound Dressing (Oasis Research)
- CollaGUARD®
- CollaMend™
- CollaSorb™
- CollaWound™
- Coll-e-derm
- Collexa®
- Collieva®
- Conexa™
- Corecyte
- Coreleader Colla-Pad
- Coretext or Protex
- CorMatrix®
- Corplex, Corplex P
- CRXa™
- Cryo-cord
- Cygnus
- Cygnus Max
- Cymetra® (Micronized AlloDerm™)
- Cytal™ (previously MatriStem®)
- Dermacyte
- Dermadapt™ Wound Dressing
- Derma-gide
- DermaPure™
- DermaSpan™

- Derm-maxx
- DressSkin
- Dermavest™
- Durepair Regeneration Matrix®
- Endoform Dermal Template™
- Epifix injectable
- *ENDUR*Agen™
- Excellagen®
- ExpressGraft™
- E-ZDerm®
- FlexiGraft®
- Floweramnioflo, Floweramniopatch
- FlowerDerm™
- Fluid flow or Fluid GF
- GammaGraft
- Geistlich Derma-Gide™
- Genesis
- GraftJacket® Xpress, injectable
- Guardian/AmniBand®
- HA Absorbent Wound Dressing
- Helicoll™
- Hyalomatrix®
- Hyalomatrix® PA
- hMatrix®
- HydraTek®
- Integra™ Bilayer Wound Matrix
- Integra® Matrix Wound Dressing (previously Avagen)
- InteguPly®
- Interfyl™
- Jaloskin®
- Kermatrix®
- Kerecis™ Omega3
- Keroxx™
- MatriDerm®
- Matrion

- MatriStem
- Matrix Collagen Wound Dressing
- Matrix HD™
- MediHoney®
- Mediskin®
- MemoDerm™
- Microderm® Biologic Wound Matrix
- MicroMatrix®
- MyOwn skin
- NeoForm™
- Neopatch or Therion
- Neox®100
- Neox®Flo
- Neox® Cord
- Neox® Wound
- Novachor
- Novafix®, Novafix DL
- NovoSorb® BTM (biodegradable temporizing matrix)
- NuCel®
- NuShield™
- Oasis® Burn Matrix
- Oasis® Ultra Tri-Layer Matrix
- Ologen™ Collagen Matrix
- Omega3 Wound (originally Merigen wound dressing)
- OrthoFlo™
- PalinGen® Flow
- PalinGen® Membrane
- PalinGen® SprotFlow
- Pelvicol®/PelviSoft®
- Permacol™
- Plurivest™
- Polycyte
- Procenta
- Progenamatrix
- ProMatrX™ ACF

- Puracol® and Puracol® Plus Collagen Wound Dressings
- PuraPly™ Wound Matrix (previously FortaDerm™)
- PuraPly™ AM (Antimicrobial Wound Matrix)
- Puros® Dermis
- RegenePro™
- ReNu™
- Repliform®
- Repriza™
- Restorigin Injectable
- Revita
- Revitalon™
- SIS Wound Dressing II
- SkinTE™
- Sport Flow™
- SS Matrix™
- Stimulen™ Collagen
- StrataGraft®
- Strattice™ (xenograft)
- Suprathel®
- Surgenex, Surfactor, and Nudyn
- Surgicord
- SurgiGRAFT™
- SurgiMend®
- Talymed®
- TenoGlide™
- TenSIX™ Acellular Dermal Matrix
- TheraForm™ Standard/Sheet
- TheraSkin®
- TissueMend
- TransCyte™
- TruSkin™
- Unite™ Biomatrix
- Veritas® Collagen Matrix
- WoundEx®, 45, 200
- WoundEx® Flow

- Woundfix, Woundfix Plus, Woundfix XPlus
- Xcellerate
- XCM Biologic® Tissue Matrix
- XenMatrix™ AB
- Xwrap.

The evidence is insufficient to determine the effects of the technology on health outcomes.

BILLING/CODING INFORMATION:

HCPCS Coding:

C9354	Acellular pericardial tissue matrix of nonhuman origin (Veritas), per sq cm (Investigational)
C9356	Tendon, porous matrix of cross-linked collagen and glycosaminoglycan matrix (TenoGlide Tendon Protector Sheet), per sq cm (Investigational)
C9358	Dermal substitute, native, nondenatured collagen, fetal bovine origin (SurgiMend Collagen Matrix), per 0.5 sq cm (Investigational)
C9360	Dermal substitute, native, nondenatured collagen, neonatal bovine origin (SurgiMend Collagen Matrix), per 0.5 sq cm (Investigational)
C9363	Skin substitute (Integra Meshed Bilayer Wound Matrix), per square cm (Investigational)
C9364	Porcine implant, Permacol, per sq cm (Investigational)
Q4100	Skin substitute, not otherwise classified
Q4101	Apligraf, per square centimeter
Q4102	Oasis Wound Matrix, per square centimeter
Q4103	Oasis Burn Matrix, per square centimeter (Investigational)
Q4104	Integra Bilayer Matrix Wound Dressing (BMWD), per square centimeter (Investigational)
Q4105	Integra dermal regeneration template (drt) or integra omnigraft dermal regeneration matrix, per square centimeter
Q4106	Dermagraft, per square centimeter
Q4107	Graftjacket, per square centimeter
Q4108	Integra Matrix , per square centimeter (Investigational)
Q4110	PriMatrix, per square centimeter
Q4111	Gammagraft, per square centimeter (Investigational)
Q4112	Cymetra, injectable, 1 cc (Investigational)
Q4113	Allograft, Graftjacket Xpress, injectable, 1 cc (Investigational)
Q4114	Integra Flowable Wound Matrix, injectable, 1 cc
Q4115	Alloskin, per square centimeter (Investigational)
Q4116	Alloderm, per square centimeter
Q4117	Hyalomatrix, per square centimeter (Investigational)
Q4118	Matristem Micromatrix, 1 MG (Investigational)
Q4121	Theraskin, per square centimeter (Investigational)
Q4122	DermACELL, DermACELL AWM or DermACELL AWM Porous, per sq cm
Q4123	AlloskinRT, per square centimeter (Investigational)
Q4124	Oasis Ultra Tri-layer wound matrix, per square centimeter (Investigational)

Q4125	Arthroflex, per square centimeter (Investigational)
Q4126	Memoderm, Dermospan, Tranzgraft or Integuply, per square centimeter (Investigational)
Q4127	Talymed, per square centimeter (Investigational)
Q4128	FlexHD, AllopatchHD, or Matrix HD, per square centimeter
Q4130	StratticeTM, per square centimeter (Investigational)
Q4132	Grafix core and grafixpl core, per square centimeter
Q4133	Grafix prime, grafixpl prime, stravix and stravixpl, per square centimeter
Q4134	hMmatrix, per square centimeter (Investigational)
Q4135	Mediskin, per square centimeter (Investigational)
Q4136	EZ-Derm, per square centimeter (Investigational)
Q4137	Amnioexcel, amnioexcel plus or biodexcel, per square centimeter (Investigational)
Q4138	BioDfence Dryflex per square centimeter (Investigational)
Q4139	AmnioMatrix or BioDMatrix, injectable, 1 cc (Investigational)
Q4140	BioDfense, per square centimeter (Investigational)
Q4141	Alloskin AC, per square centimeter (Investigational)
Q4142	XCM Biologic Tissue Matrix, per square centimeter (Investigational)
Q4143	Repriza, per square centimeter (Investigational)
Q4145	Epifix, injectable, 1 mg (Investigational)
Q4146	TenSIX, per square centimeter (Investigational)
Q4147	Architect, Architect PX, or Architect FX, extracellular matrix, per square centimeter (Investigational)
Q4148	Neox cord 1k, neox cord rt, or clarix cord 1k, per square centimeter (Investigational)
Q4149	Excellagen, 0.1 cc (Investigational)
Q4150	Allowrap DS or dry, per square centimeter (Investigational)
Q4151	Amnioband or guardian, per square centimeter
Q4152	Dermapure, per square centimeter (Investigational)
Q4153	Dermavest and plurivest, per square centimeter (Investigational)
Q4154	Biovance, per square centimeter
Q4155	Neoxflo or clarixflo, 1 mg (Investigational)
Q4156	Neox 100 or clarix 100, per square centimeter (Investigational)
Q4157	Revitalon, per square centimeter (Investigational)
Q4158	Kerecis omega3, per square centimeter (Investigational)
Q4159	Affinity, per square centimeter
Q4160	Nushield, per square centimeter (Investigational)
Q4161	Bio-connekt wound matrix, per square centimeter (Investigational)
Q4162	Woundex flow, bioskin flow, 0.5 cc (Investigational)
Q4163	Woundex, bioskin, per square centimeter (Investigational)
Q4164	Helicoll, per square centimeter (Investigational)
Q4165	Keramatrix or kerasorb, per square centimeter (Investigational)
Q4166	Cytal, per square centimeter (Investigational)
Q4167	Truskin, per square centimeter (Investigational)
Q4168	Amnioband, 1 mg (Investigational)
Q4169	Artacent wound, per square centimeter (Investigational)
Q4170	Cygnus, per square centimeter (Investigational)

Q4171	Interfyl, 1 mg (Investigational)
Q4173	Palingen or palingen xplus, per square centimeter (Investigational)
Q4174	Palingen or promatrix, 0.36 mg per 0.25 cc (Investigational)
Q4175	Miroderm, per square centimeter (Investigational)
Q4176	Neopatch or Therion, per square centimeter (Investigational)
Q4177	Floweramnioflo, 0.1 cc (Investigational)
Q4178	Floweramniopatch, per square centimeter (Investigational)
Q4179	Flowerderm, per square centimeter (Investigational)
Q4180	Revita, per square centimeter (Investigational)
Q4181	Amnio wound, per square centimeter (Investigational)
Q4182	Transcyte, per square centimeter (Investigational)
Q4183	Surgigraft, per square centimeter (Investigational)
Q4184	Cellesta or cellesta duo, per square centimeter (Investigational)
Q4185	Cellesta flowable amnion (25 mg per cc); per 0.5 cc (Investigational)
Q4186	Epifix, per square centimeter
Q4187	Epicord, per square centimeter
Q4188	Amnioarmor, per square centimeter (Investigational)
Q4189	Artacent ac, 1 mg (Investigational)
Q4190	Artacent ac, per square centimeter (Investigational)
Q4191	Restorigin, per square centimeter (Investigational)
Q4192	Restorigin, 1 cc (Investigational)
Q4193	Coll-e-derm, per square centimeter (Investigational)
Q4194	Novachor, per square centimeter (Investigational)
Q4195	Puraply, per square centimeter (Investigational)
Q4196	Puraply am, per square centimeter (Investigational)
Q4197	Puraply xt, per square centimeter (Investigational)
Q4198	Genesis amniotic membrane, per square centimeter (Investigational)
Q4200	Skin te, per square centimeter (Investigational)
Q4201	Matrion, per square centimeter (Investigational)
Q4202	Kerxxx (2.5g/cc), 1cc (Investigational)
Q4203	Derma-gide, per square centimeter (Investigational)
Q4204	Xwrap, per square centimeter (Investigational)
Q4205	Membrane graft or membrane wrap, per square centimeter (Investigational)
Q4206	Fluid flow or fluid GF, 1 cc (Investigational)
Q4208	Novafix, per square centimeter (Investigational)
Q4209	Surgraft, per square centimeter (Investigational)
Q4210	Axolotl graft or axolotl dualgraft, per square centimeter (Investigational)
Q4211	Amnion bio or Axobiomembrane, per square centimeter (Investigational)
Q4212	Allogen, per cc (Investigational)
Q4213	Ascent, 0.5 mg (Investigational)
Q4214	Cellesta cord, per square centimeter (Investigational)
Q4215	Axolotl ambient or axolotl cryo, 0.1 mg (Investigational)
Q4216	Artacent cord, per square centimeter (Investigational)
Q4217	Woundfix, BioWound, Woundfix Plus, BioWound Plus, Woundfix Xplus or BioWound Xplus, per square centimeter (Investigational)
Q4218	Surgicord, per square centimeter (Investigational)
Q4219	Surgigraft-dual, per square centimeter (Investigational)

Q4220	BellaCell HD or Surederm, per square centimeter (Investigational)
Q4221	Amniowrap2, per square centimeter (Investigational)
Q4222	Progenamatrix, per square centimeter (Investigational)
Q4226	MyOwn skin, includes harvesting and preparation procedures, per square centimeter (Investigational)
Q4227	Amniocore, per square centimeter (Investigational)
Q4229	Cogenex amniotic membrane, per square centimeter (Investigational)
Q4230	Cogenex flowable amnion, per 0.5 cc (Investigational)
Q4231	Corplex p, per cc (Investigational)
Q4232	Corplex, per square centimeter (Investigational)
Q4233	Surfactor or nudyn, per 0.5 cc (Investigational)
Q4234	Xcellerate, per square centimeter (Investigational)
Q4235	Amniorepair or altiely, per square centimeter (Investigational)
Q4237	Cryo-cord, per square centimeter (Investigational)
Q4238	Derm-maxx, per square centimeter (Investigational)
Q4239	Amnio-maxx or amnio-maxx lite, per square centimeter (Investigational)
Q4240	Corecyte, for topical use only, per 0.5 cc (Investigational)
Q4241	Polycyte, for topical use only, per 0.5 cc (Investigational)
Q4242	Amniocyte plus, per 0.5 cc (Investigational)
Q4244	Procenta, per 200 mg (Investigational)
Q4245	Amniotext, per cc (Investigational)
Q4246	Coretext or protext, per cc (Investigational)
Q4247	Amniotext patch, per square centimeter (Investigational)
Q4248	Dermacyte amniotic membrane allograft, per square centimeter (Investigational)
Q4249	Amniplay, for topical use only, per square centimeter (Investigational)
Q4250	Amnioamp-mp, per square centimeter (Investigational)
Q4251	Vim, per square centimeter (Investigational)
Q4252	Vendaje, per square centimeter (Investigational)
Q4253	Zenith amniotic membrane, per square centimeter (Investigational)
Q4254	Novafix dl, per square centimeter (Investigational)
Q4255	Reguard, for topical use only, per square centimeter (Investigational)

ICD-10 Diagnosis Codes That Support Medical Necessity:

C50.011 – C50.019 C50.111 – C50.119 C50.211 – C50.219 C50.311 – C50.319 C50.411 – C50.519 C50.611 – C50.619 C50.811 – C50.819 C50.911 – C50.919	Malignant neoplasm of breast
D05.00 – D05.92	Carcinoma in situ of breast
E08.621	Diabetes mellitus due to underlying condition with foot ulcer
E08.622	Diabetes mellitus due to underlying condition with other skin ulcer
E09.621	Drug or chemical induced diabetes mellitus with foot ulcer
E09.622	Drug or chemical induced diabetes mellitus with other skin ulcer
E10.621	Type 1 diabetes mellitus with foot ulcer

E10.622	Type 1 diabetes mellitus with other skin ulcer
E11.621	Type 2 diabetes mellitus with foot ulcer
E11.622	Type 2 diabetes mellitus with other skin ulcer
E13.621	Other specified diabetes mellitus with foot ulcer
E13.622	Other specified diabetes mellitus with other skin ulcer
I83.001 – I83.029	Varicose veins of lower extremities with ulcer
I83.201 – I83.229	Varicose veins of lower extremities with both ulcer and inflammation
Q81.2	Epidermolysis bullosa dystrophica
T20.20xA – T20.39xS T20.60xA – T20.79xS	Burn and corrosion of head, face and neck
T21.20xA – T21.39xS T21.60xA – T21.79xS	Burn and corrosion of trunk
T22.20xA – T22.399S T22.60xA – T22.799S	Burn and corrosion of shoulder and upper limb except wrist and hand
T23.201A – T23.399S T23.601A – T23.799S	Burn and corrosion of wrist and hand
T24.201A – T24.399S T24.601A – T24.799S	Burn and corrosion of lower limb, except ankle and foot
T25.211A – T25.399S T25.611A – T25.799S	Burn and corrosion of ankle and foot
T30.0 – T32.99	Burns classified according to extent of body surface involvement
T34.011A – T34.99XS	Frostbite with tissue necrosis

REIMBURSEMENT INFORMATION:

Apligraf® is limited to five (5) applications per ulcer. The safety and the effectiveness of Apligraf have not been established for patients receiving greater than 5 applications.

Dermagraft® is limited to eight (8) applications per treatment site over a twelve (12) week period. Dermagraft has not been studied in patients receiving greater than 8 device applications.

OrCel™ is limited to a single, one-time application per donor site. No more than eight (8) pieces should be used per donor site. The safety and effectiveness of OrCel has not been evaluated in burn patients with split thickness donor sites larger than 288cm (8 pieces of OrCel).

Safety and effectiveness of **re-treatment** of a single wound using Apligraf®, Dermagraft® or OrCel™ has not been established and is considered **experimental or Investigational**.

PROGRAM EXCEPTIONS:

Federal Employee Program (FEP): Follow FEP guidelines.

State Account Organization (SAO): Follow SAO guidelines.

Medicare Advantage products:

The following National Coverage Determination (NCD) was reviewed on the last guideline reviewed date: Porcine skin and Gradient Pressure Dressing (270.5) located at cms.gov.

The following Local Coverage Determination (LCD) was reviewed on the last guideline reviewed date: Application of Skin Substitute Grafts for Treatment of DFU and VLU of Lower Extremities (L36377) located at fcso.com.

DEFINITIONS:

Nonhealing- less than a 20% decrease in wound area with standard wound care for at least 2 weeks.

RELATED GUIDELINES:

[Amniotic Membrane and Limbal Stem Cell Transplantation for the Treatment of Ocular Conditions, 02-65000-19](#)

OTHER:

None.

REFERENCES:

1. Agency for Healthcare Research and Quality (AHRQ), Negative Pressure Wound Therapy Devices, Technology Assessment Report, 2009. Accessed at ahrq.gov 08/02/12.
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COMMITTEE APPROVAL:

This Medical Coverage Guideline (MCG) was approved by the Florida Blue Medical Policy & Coverage Committee on 03/25/21.

GUIDELINE UPDATE INFORMATION:

01/01/01	New Medical Coverage Guideline.
01/01/02	Annual HCPCS coding update.
01/01/03	Annual HCPCS coding update.
02/15/03	Annual Review.
06/15/04	Unscheduled review and revision to guideline; consisting of updated references, deleted J7350 and changed name of MCG from Apligraf (Graftskin) to Human Skin Equivalent Grafts.
01/01/05	Annual HCPCS coding update: consisting of addition of J7343 and J7344 and deletion of Q0182 and Q0183.
01/01/06	Annual HCPCS coding update: added 15000, 15340, 15341, 15360, 15361, 15365, and 15366. Deleted: 15342, 15343, J7343. Revised: J7340, J7342, and J7344.
08/15/06	Biennial review; new information added for Integra, TransCyte, allograft, OrCel, xenograft, AlloDerm; revision to code information; code J7343 added.
01/01/07	Annual HCPCS coding update: added J7345, J7346; deleted 15000 – 15400 & changed to 15002 – 15005 to describe codes used for graft site preparation & graft application.
03/15/07	Revision to guideline; consisting of addition of OASIS Wound Matrix (J7341), revision to criteria for Apligraf (J7340), revision to code information for J7344 and J7343, updated references.
06/15/07	Reformatted guideline.

08/15/07	Review, coverage statements maintained, references updated.
01/01/08	Annual HCPCS coding update: deleted J7345; added J7347, J7348, J7349.
01/01/09	Annual HCPCS coding update: added HCPCS codes Q4100 – Q4114; updated ICD-9 codes 707.10 – 707.19 & 707.8; and deleted HCPCS codes J7340-J7349.
07/01/09	Biennial review: description section, Alloderm position statement, coding and references updated. HCPCS 3 rd quarter coding update: added new codes Q4115 and Q4116.
10/15/10	Revision; related ICD-10 codes added.
01/01/11	Annual HCPCS coding update. Added Q4117 – Q4121; revised Q4101 – Q4116; deleted Q4109.
07/15/11	Scheduled review; position statements maintained, coding section and references updated.
01/01/12	Annual HCPCS coding update; added HCPCS codes Q4122-Q4130; CPT coding section updated.
07/15/12	Annual review; position statements, billing/coding information, description section, Medicare program exception, and references updated; formatting changes.
01/01/13	Annual HCPCS update; added Q4131-Q4136; revised Q4119, Q4126, & Q4128. Experimental list updated; formatting changes.
07/15/13	Annual review; description section, position statement, and references updated; formatting changes.
01/01/14	Annual HCPCS update. Added codes Q4137-Q4149. Position statement updated; formatting changes.
03/15/14	Annual review; position statements, Coding, Description, and references updated; formatting changes.
06/15/14	Revision; references updated.
01/01/15	Annual HCPCS/CPT update. Added codes Q4150-Q4160; revised codes Q4119 & Q4147.
03/15/15	Annual review; description and position statement section updated; coding and references updated; formatting changes.
10/01/15	Revision; ICD9 and ICD10 coding sections updated.
11/01/15	Revision: ICD-9 Codes deleted.
01/01/16	Annual HCPCS/CPT update; codes Q4161-Q4165 added; code Q4153 revised.
08/15/16	Revision; policy title, description section, position statement section, coding, and references updated; formatting changes.
10/01/16	Revision; formatting changes. Investigational product list updated.
11/15/16	Revision; coding section updated.
01/01/17	Annual CPT/HCPCS update. Added Q4166-Q4175; revised Q4105 & Q4131; deleted Q4119, Q4120, Q4129.
03/15/17	Revision; Position statements including treatment of diabetic ulcers, amniotic membrane, and investigational product list updated; Code Q4151, program exception, and references updated.
04/15/17	Revision; code C9349 deleted.
08/15/17	Revision; Integra Omnigraft deleted from investigational product list and added to bullet for Integra® Dermal Regeneration Template.
01/01/18	Annual CPT/HCPCS update. Added codes Q4176-Q4182; revised codes Q4132, Q4133, Q4148, Q4156, Q4158, Q4162, Q4163.
04/15/18	Revision; description, position statements, coding, and references updated.
01/01/19	Annual CPT/HCPCS coding update. Added codes Q4183-Q4204; revised codes Q4133 & Q4137; deleted codes Q4131 & Q4172.
04/15/19	Review; Investigational product list and references updated.

07/01/19	Revision; Investigational product list updated.
10/01/19	Quarterly CPT/HCPCS coding update. Added codes Q4205-Q4226; revised codes Q4122, Q4165, Q4184.
04/15/20	Review; position statements and references updated.
07/01/20	Quarterly CPT/HCPCS coding update. Added codes Q4227-Q4248; revised code Q4176.
08/15/20	Revision; coding section updated.
10/01/20	Quarterly CPT/HCPCS coding update. Added codes Q4249, Q4250, Q4254, Q4255.
04/15/21	Review; Position statements, investigational product list, coding and references updated.
05/21/21	Revision; investigational product list updated.
10/01/21	Quarterly CPT/HCPCS coding update. Codes Q4251-Q4253 added; codes Q4228 & Q4236 deleted.