

HELICOLL® COMPARED TO OTHER PRODUCTS

HELICOLL® COMPARISON WITH OTHER FDA APPROVED PRIME PRODUCTS:

https://helicoll.com/wp-content/uploads/2024/03/Helicoll_comparison_with_other_FDA_approved_prime_products_table.pdf

Unique Benefits to Safety:

All of the following aspects are well addressed with proper production and management documents to comply the Medical Product regulations.

- (i) Periodic monitoring of the results of the risk analysis
- (ii) Specification of materials, and manufacturing/special processing
- (iii) Specifications, drawings and circuit diagrams for components, sub- assemblies and the complete product including packaging, where appropriate.
- (iv) The specifications of the checks, tests and trials that are intended to be carried out as part of routine production
- (v) The performances and compatibilities intended by the manufacturer
- (vi) Labeling, including any instructions for use
- (vii) Identification of 'shelf-life' reflected by any 'use by' date, or other 'lifetime' of the device(s)
- (viii) Results of Bench Testing
- (viii) Clinical data
- (ix) Post market surveillance
- (x) Documentation and reporting Design Changes

Additional Non-clinical Benefits:

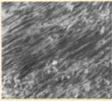
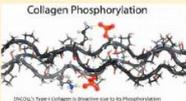
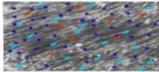
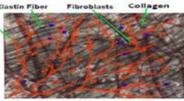
HELICOLL is a better cost-effective product than most other advanced skin substitutes on the market. Also, the easy use, storage and long shelf life are all added features to the product.

Our team of Wound Care specialists are extremely happy to assist your team of experts to bring the public awareness to the regenerative aspects of wound care that uses nanotechnology that bio-mimics the physiological biochemical pathways of wound healing process.

Names of close Competitors:

Q4101 Apligraf	Q4134 hMatrix
Q4102 Oasis Wound Matrix	Q4135 Mediskin
Q4104 Integra BMWD	Q4136 Ezderm
Q4106 Dermagraft	Q4150 Allowrap DS or Dry 1 sq cm
Q4107 GraftJacket	Q4151* AmnioBand, Guardian 1 sq cm
Q4110 Primatrix	Q4152* Dermapure 1 square cm
Q4111 Gammagraft	Q4153 Dermavest 1 square cm
Q4115 Alloskin	Q4154* Biovance 1 square cm
Q4116 Alloderm	Q4160 NuShield 1 square cm
Q4121 Theraskin	Q4161 Bio-Connekt per square cm
Q4122 Dermacell, awm, porous sq cm	Q4163 Woundex, bioskin, per sq cm
	Q4164 Helicoll, per sq cm
	Q4165 Keramatrix, kerasorb sq cm

Q4123 Alloskin	Q4186 Epifix
Q4124 Oasis Tri-layer Wound Matrix	Q4195+ Puraply 1 sq cm
Q4126 Memoderm/derma/tranz/integup	Q4203 Derma-gide, 1 sq cm
Q4127 Tallymed	Q4204 Xwrap 1 sq cm
Q4133 Grafix stravix prime pl sq cm	Q4221 Amniowrap2 per sq cm

117 MEDICARE-APPROVED SKIN SUBSTITUTES: PROPERTIES BY PRODUCT CATEGORY				
PRODUCT ORIGIN	PRODUCT TYPE	PRODUCT DESCRIPTION	# of Total	PRODUCT CLINICAL PROPERTIES
XENOGRAFT (ANIMAL ORIGIN)	EXTRACTED COLLAGEN SHEET	<p>Helicoll</p> <p>Patented Highly Purified, bioactive, non-immunogenic type-I collagen product. Helicoll's unique collagen scaffold with approx. 20µ Porosity and native staggered fibres.</p>   <p>Collagen Phosphorylation</p> <p><small>ENCOLL's Type-I Collagen is bioactive due to its Phosphorylation</small></p>	1	  <p>Helicoll Bioactive type-I collagen attracts cells & factors in 1 to 3 days</p> <p>Tissue regeneration in 4 to 5 days, not shown with other products</p> <ul style="list-style-type: none"> Highly regenerative, phosphorylated, clinically effective, Type-I collagen.
		<p>Primatrix, Puraply, Integra & GraftJacket</p> <p>Partially pure collagen mostly cross-linked with non-native, random distribution of fibres.</p> 	12	<ul style="list-style-type: none"> Purification not patented. Random, non-native structural configuration. Freeze-drying results in matrix with average porosity > 800µ that is not conducive for ideal cell infiltration. Gamma sterilization is not ideal as it may denature and cross-link the protein resulting in lesser bioactivity.
	<p>Oasis</p> <p>Porcine Intestinal Wall</p> <p>Cyral Wound Matrix</p> <p>Porcine urinary bladder matrix</p> <p>Mediskin</p> <p>Porcine dermal and epidermal layer</p> <p>Kerecis Omega3 Wound</p> <p>Fish Skin</p>	7	<ul style="list-style-type: none"> All intact tissue membrane products, by default, contain immunogenic components like Elastin (approximately > 15% w/w) and Type-III Collagen etc. In case of Kerecis, chitosan is an additional contaminant. Kerecis with Omega3 Fatty acid may result in production of free radicals that damages tissue cells resulting in lesser regeneration. 	
ALLOGRAFT (HUMAN ORIGIN)	INTACT TISSUE DERIVED MEMBRANE	<p>Alloskin, Alloderm, Dermagraft, Truskin, Miroderm, Progenamatrix</p> <p>Human Skin</p>	21	<ul style="list-style-type: none"> Direct or indirect cross-linking done to reduce their immunogenicity may constrain bioactivity. Restricted bio-effectivity and biological regenerative capabilities in the host tissue.
		<p>Amnion Bio, Biovance, Epifix, Restorigin, Woundfix Biowound, Xwrap</p> <p>Amnion</p> <p>Cellesta Cord & Surgicord</p> <p>Umbilical Cord</p> <p>Grafix, NuShield, Neogatch,</p> <p>Placenta</p>	66	
		<p>Apligraf & Dermagraft</p> <p>Cell Seeded & Others</p>	10	<ul style="list-style-type: none"> Impaired bioactivity and biocompatibility.

What Differentiates Encoll's Helicoll product from the Competitors:

Encoll has over thirty years expertise in the field of surgical applications of collagen. The founder and CEO of Encoll has technical contributions in the development of standards for manufacturing surgical grade collagen. This fact is evidenced by Dr. Gunasekaran's active involvement in the steering committee along with the FDA group headed by Dr. David Kaplan and other experts at ASTM (American Standards for Test Methods). Resulted in 1995 a ASTM as the guidelines to the manufacturers of Type-I collagen. (Ref: ASTM: F 2212-09 Standard Guide for Characterization of Type I Collagen as Starting Material for Surgical Implants and Substrates for Tissue Engineered Medical Products).

Encoll's expertise in the technical aspects of the development of non-immunogenic biological construct is enormous. Among the other fifty products in the market that are Medicare approved, almost half of them are manufactured from intact tissues. The manufacturers of such products are given authority to claim their products are "Biological skin substitutes". Unfortunately, these manufacturers don't realize that 15% elastin is still present in their final product which is NOT bio-compatible. Thereby, they have to cross-link the finished product to eliminate the rejection of those products. This action results in the loss of bio-activity for the entire product. This is the MAJOR difference when compared to Helicoll that is not endpoint cross-linked in any manner.

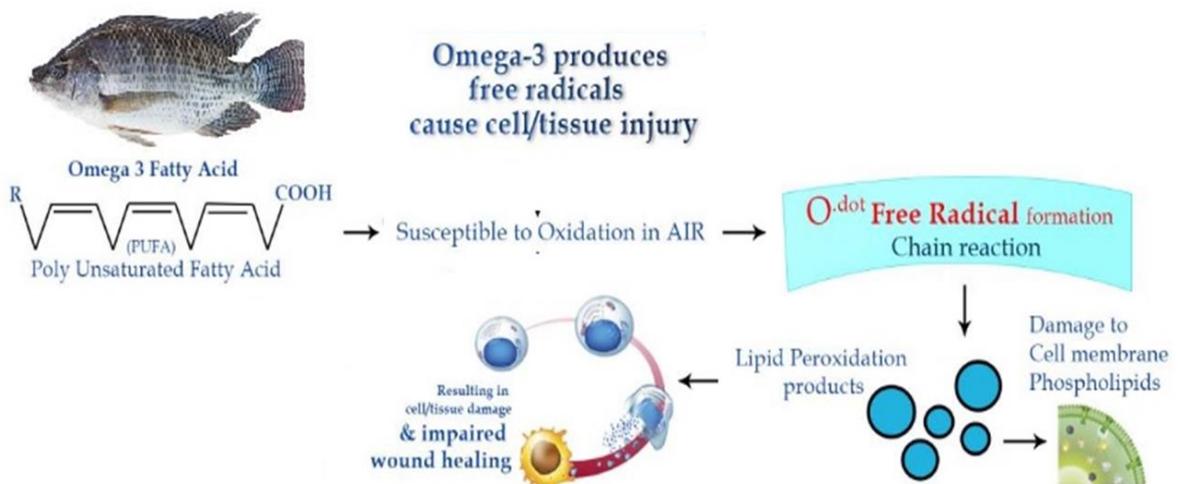
ALLOGRAFT		XENOGRAFT	
Grafix  PLACENTA	 AMNION	Oasis  INTESTINAL MUCOSA	 URINARY BLADDER
Epifix XWrap		Cytal	
<p>Intact tissue membrane products, by default contain immunogenic components like Elastin (approximately 15% w/w) and Type-III Collagen etc.</p> <p>To reduce their immunogenicity, every product is directly or indirectly crosslinked which constrains the natural bioactivity of Type-I collagen.</p> <p>Such restricted bio-effectivity of the Type-I collagen affects its natural biological regenerative capabilities in the host tissue.</p>			

Another set of products fall under the allograft category. The same hypothesis applies to these products where they produce an inferior outcome.

Another set of manufacturers who produce products with cultured cells are also having difficulties to manage the incompatibility or immunogenicity of certain structural proteins synthesized by those cells harvested from another individual. Furthermore, all such cell seeded products have an extremely high risk in maintaining sterility of their end product with a very short shelf life and an enormous cost of production.

With respect to Piscean collagen, even the type-I collagen of a fish is not equivalent to that of human type-I collagen. It may not be biocompatible, as its amino acid sequence is not matching with that of a human type-I collagen (only 77.9% homology). On the other hand, Helicoll bovine type-I collagen is more biocompatible as it is 97.4% homologous to the human type-I collagen.

Piscean collagen also may have contaminating chitin (chitosan) possibly due to the remnants of scales in fish skin. Besides, the presence of Omega 3 fatty acids is more damaging to the tissues than its benefits. Fatty acids when stored in atmospheric air would release free radicals to damage the adjacent tissues.



I prefer the reviewer of this document to consider the pros and cons of intact tissue based skin substitutes. Naturally all intact tissue based products do have at least 15% of high immunogenic compound namely Elastin, besides other allergenic biological molecules like glycosaminoglycans and certain types of collagen other than type-I collagen.

Both the above products containing potential immunogenic molecules have to be surface charge modified (cross-linked) to minimize their immunogenicity. As a result, the above products are not capable to be a bio-active construct for faster healing and repair of the damaged tissues.



Other collagen preparations (ex. **Puraply, Primatrix and Integra**) as a result of lyophilization, yield random structural configuration that is non-native with avg. porosity > 800 μ which is non-conducive for cell infiltration. Its bioactivity stays behind the bioactivity of sedimental preparation of **HELICOLL** type-I collagen that yields native parallel fibrils with porosity of approx. 20 μ to attract more cells/regenerative factors. Additionally, **Helicoll** is EtO gas sterilized that does not denature the protein unlike gamma sterilization which is used by most other collagen preparations. Collagen when cross-linked, the natural bioactivity may be significantly minimized.

HELICOLL:

On the other hand, Helicoll is nothing but highly bioactive and bio-compatible type-I collagen construct. That's why, Helicoll has been clinically documented for faster wound healing. (see the recent clinical study publications: <https://helicoll.com/clinical-studies/>) It encourages the formation of **new blood capillaries within 4 to 5 days** upon the application of the product over the wound.

No other skin substitute has shown such an advanced clinical result.