Using the Matrix: Efficacy of Extracellular Matrix in Treatment Algorithm for Recurrent Benign Esophageal Stricture Therapy

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Background

- Recurrent benign esophageal stricture (RBES) is caused by chronic inflammation from GERD, radiation, or surgery.
- Despite various therapeutic options including dilation, incision, steroid injection, and self-expandable metal stents, efficacy of these interventions remains suboptimal ranging between 20% to 63.6%.
- AIM: Propose and design a new treatment modality using an extracellular matrix (ECM) in the algorithm of treating RBES to extend duration of time and decrease frequency of treatments.

Methods

- We designed a proof-of-concept endoscopic technique that utilizes a biodegradable scaffold membrane composed of ECM as part of our stricture management algorithm to stimulate site specific tissue repair.
- Ten patients, mean age 61 years (38-80) with 6 females:4 males, presented with four types of RBES (40% peptic, 30% anastomotic, 20% post Zenker's septotomy and 10% fistula-related) were recruited.
- Mean follow up duration was 811 days (42-1726).
- Patients underwent endoscopy with placement of ECM and clip placement or stent placement with follow up EGD to ensure correct placement
- Other modalities used: Steroid injection, balloon dilation, incisional therapy, and esophageal stenting

	Average Pre- ECM Treatment Frequency (Weeks)	Average Post- ECM treatment Frequency (Weeks)	Average Percent Reduction in Treatment Frequency	Average Percent Reduction in Mean number of Procedures	Other Modalities used
Post Zenker's Septotomy (n=2)	14.00	N/A	N/A	N/A	Flexible endoscopic incisional therapy
Peptic Stricture (n=4)	6.50	9.33	17%	30%	Incision, balloon dilation, steroid Injection, Fully covered metal esophageal stent
Anastomotic Stricture (n=3)	4.67	5.00	0%	38%	Incision, Balloon dilation, steroid Injection
Fistula (n=1)	4.00	4.00	0%	N/A	Sealant and fully covered metal esophageal stent

Table 1.

- Post Zenker's septotomy strictures, there
 was 100% complete resolution of
 symptoms at two months with no further
 interventions required.
- There was no change in therapy frequency in patients with post-surgical anastomotic esophageal strictures or fistula-related stenosis.
- In peptic strictures, the average therapy interval was extended from 6.5 weeks to 9.3 weeks after ECM therapy, leading to 17% reduction in treatment frequency following ECM.
- The mean number of procedures required was reduced by 30% for peptic strictures and by 38% for anastomotic strictures.

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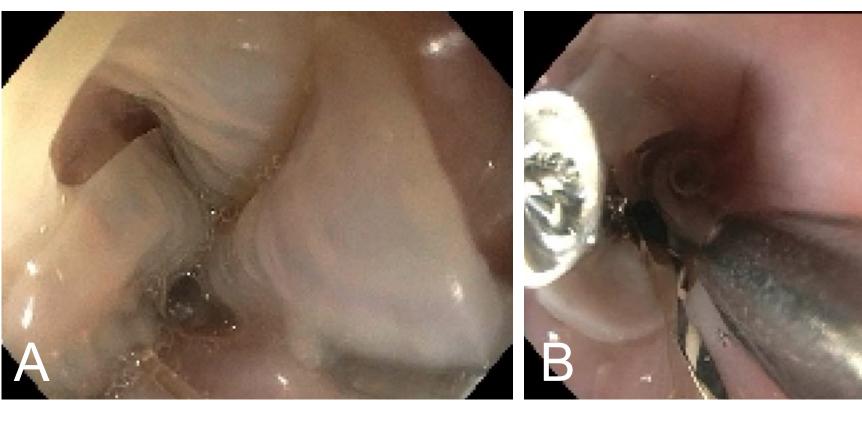


Figure 1. (A) ECM unfurling in esophagus. (B) ECM membrane being clipped into position.

Discussion

- Proposed pathophysiology: ECM degradation triggers chemotactic and mitogenic activity for progenitor cells and inhibit proliferation of differentiated endothelial cells slowing down the process of recurrence
- Our novel technique demonstrates that the utilization of ECM membrane as part of the treatment algorithm of RBES
 - reduces both the frequency and
 - the number of repeat interventions in select patients.
- Post septotomy and peptic strictures responded the best to ECM therapy in conjunction with other modalities compared to anastomotic strictures.
- Further larger prospective studies are required for increased understanding of ECM therapy response in various disease states.

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