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CASE REPORT

Fine needle aspiration cytology as a preliminary diagnostic tool in chondroid syringoma: a case report and review

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Abstract: We report a case of chondroid syringoma (CS) in a 44-year-old male. He presented with a firm asymptomatic nodule in his left upper lip of 2-year duration. The initial clue to the diagnosis was made on fine needle aspiration cytology (FNAC), and a final diagnosis was based on histopathological examination. The case highlights the importance of FNAC in providing clues to the diagnosis of suspected cases of chondroid syringoma before performing large excisions and repair, which would require more skill and time. We have also reviewed the cytological findings of all the cases of benign CS reported until the current date.

Keywords: chondroid syringoma, pleomorphic adenoma, adnexal tumours, fine needle aspiration cytology

Introduction

Chondroid syringoma (CS) is a rare cutaneous tumour originating from eccrine and apocrine sweat glands with both epithelial and mesenchymal components. The incidence is low, forming less than 0.01% of primary cutaneous tumours.¹ The clinical diagnosis is challenging. Histopathology is imperative to reach the diagnosis. The role of fine needle aspiration cytology (FNAC) has been underused in the past and is infrequently utilized for diagnosis or preoperative assessment. We report a case of benign chondroid syringoma of the upper lip, initially diagnosed by FNAC and later confirmed by histopathology. A literature review on the topic has also been done. The key words "chondroid syringoma" and "pleomorphic adenoma" were used to search databases which included PubMed, Google Scholar, Cochrane library and Hinari, and relevant papers were retrieved.

Case presentation

A 44-year-old male presented with an asymptomatic, progressive swelling over the left half of his upper lip of 2-years duration. The swelling was initially small and had gradually increased to present dimensions. There was no history of trauma, discharge or any other similar lesion in the body. There was no history of symptoms suggestive of systemic illnesses. On examination, there was a firm, non-fluctuant, skin coloured, non-tender nodule of size 1.5 cm×1.5 cm with overlying normal skin (Figure 1). The nodule was fixed to the skin, but freely mobile over underlying structures. There was no regional lymphadenopathy.



Figure I A 1.5 cm diameter nodule on left half of upper lip before excison.

FNAC of the lesion was done which showed aggregates, acini and singly scattered benign epithelial cells along with myoepithelial cells and chondromyxoid stromal fragments



Figure 2 FNAC: aggregates, acini and single scattered benign epithelial cells along with myoepithelial cells and chondromyxoid stromal fragments (Giemsa stain, ×40). Abbreviations: FNAC, fine needle aspiration cytology.

(Figure 2). Epithelial cells were round to polygonal with basophilic dense moderate cytoplasm and central to eccentric, round to oval nuclei with bland chromatin on a background of myxoid material, thus pointing towards the possibility of chondroid syringoma (Figure 3A and B).

The nodule was excised and the whole specimen was sent for histopathological examination. The cut section showed homogenous grey white areas. The hematoxylin-eosin stain revealed cystic structures with cystically dilated ducts, nests and glandular structures lined by bland looking epithelial cells along with surrounding chondromyxoid stroma (Figure 4A and B). Histopathology confirmed the cytological diagnosis of CS. Immunohistochemistry could not be done because of unavailability in the centre. Excision site was healthy during the postoperative period and no recurrence was observed after 6 months of follow-up.

Discussion

CS is a benign cutaneous tumor with male preponderance (male to female ratio of 5:8),^{2–5} and is predominantly seen on head and neck regions with predilection for upper lips, nose and cheeks. Rare sites on the face include the orbit, eyelids, and medial canthus.^{6–9} Other uncommon sites are the back, axilla, thighs, extremities and genitalia.^{2,3,5} The tumor presents as an asymptomatic, solitary, skin coloured, firm, and non-tender slow growing nodule. A tumor in the orbit may lead to exophthalmos.⁸ The size ranges from 0.3 cm to 3 cm. Nodules exceeding 5 cm¹⁰ and 10 cm^{11,12} in diameter have also been reported. The clinical profiles of CS from five large retrospective studies are compared in Table 1.



Figure 3 (A) FNAC smear showing sheet of myoepithelial cells with basophilic dense cytoplasm and central to eccentric, round to oval nuclei with bland chromatin (Giemsa stain, ×200). (B) Cluster of epithelial cells with scattered myoepithelial cells in a chondromyxoid background (Giemsa stain, ×200). Abbreviations: FNAC, fine needle aspiration cytology.



Figure 4 (A) Section showing tumor composed of ducts and glandular structures lined by bland looking epithelial and myoepithelial cells with surrounding chondromyxoid stroma (H&E stain, ×40). (B) Focal areas showing ossification and keratinous cyst filled with keratin (H&E stain, ×40). (B) Focal areas showing ossification and keratinous cyst filled with keratin (H&E stain, ×40). (B) Focal areas showing ossification and keratinous cyst filled with keratin (H&E stain, ×40).

Because of its rarity, asymptomatic and subcutaneous nature, the clinical diagnosis of this condition is often missed as evidenced by the diagnoses made before histopathological examination revealed in Table 2. The differential diagnosis of such presentations include dermoid or sebaceous cyst, pilar cyst, calcifying epithelioma, or a solitary trichoepithelioma, dermatofibroma, lymph node, hamartoma, basal cell carcinoma, and seborrheic keratosis,⁴ with no role of non-invasive investigations like X-ray,^{14,15} ultrasonography,^{11,15,16} MRI,¹⁰ or CT scan⁸ in the diagnosis. FNAC and biopsy so far remains the gold standard for the diagnosis. FNAC which is easy to perform, is established in the literature for making early diagnosis of CS and is reviewed in Tables 2 and 3.

The origin of CS is from both secretory and ductal segments of eccrine or apocrine sweat glands. It is a mixed tumor with epithelial and mesenchymal components and resembles the pleomorphic adenoma of salivary glands. The first attempt to diagnose and document CS on FNAC was made in 1988 by Masood et al.¹⁷ The aspirate can be thick, mucoid and sometimes gelatinous with moderate cellularity. Thin aspirate may be associated with scanty stromal elements.¹⁸ The mucoid material stains positive with alcian blue and mucicarmine.^{17,19} The epithelial cells arrangement is highly variable. The cells can appear singly, scattered, in groups or as sheets, attached either loosely or cohesively.^{12,14,17,20-22} As in our case, acini^{9,19} and papillary^{16,23} configurations have also been noted. The individual cells are small to medium sized, well-defined, monomorphic, round-to-oval-to-ovoid-to-polygonal, with moderate to dense cytoplasm. The cytoplasm can be eosinophil to amphophilic, imparting a plasmacytoid appearance.²⁰ The nuclei are small, monomorphic, round, oval, ovoid or elongated, central to eccentric in location with fine, evenly distributed chromatin.^{15,17,20,21,24} Nuclear atypia is a rare finding without any propensity to develop into malignancy.²⁵ Anisonucleosis, conspicuous nucleoli or nuclei with clear halos may be suggestive of neoplastic changes, but malignancy can be safely ruled out in the absence of other features.¹⁴ The background is chondroid,^{17,22} myxoid,^{19,20,24} or chondromyxoid^{15,16,21,26-28} which can be scant¹⁸ to abundant.²⁹ Myoepithelial cells also appear in clusters or aggregates, dispersed along with epithelial cells in the stroma, and give plasmacytoid appearance with dark nuclei.^{21,23,29} Macrophages are uncommonly seen and have been reported along with cystic changes by Khan et al.¹⁸ In our case, foamy macrophages were evident on a myxoid background but without any cystic changes. Immunostaining differentiates the two components, as epithelial membrane antigen (EMA) and cytokeratin stains the epithelial cells, while S-100 makes the myoepithelial part evident.^{14,17}

Cytology in addition to clinical features like site and size, can be a tool to differentiate benign from malignant CS.³⁰ Rarely, a benign tumor may turn aggressive and go into malignant phase.³¹ Poor prognosis, metastasis and recurrences following excision are attributes of malignant CS.32-35 The clinical features that differentiates it from benign CS include female preponderance, predilection for extremities and size exceeding 3 cm.^{17,30,36,37} Studies describing FNAC findings of malignant CS are also scarce. In 1997, Mishra et al³⁶ made the first conclusive diagnosis of malignant CS on FNAC. Haemorrhagic aspirate, hypercellularity, pleomorphic epithelium, dyshesiveness of cells, intranuclear and intracytoplasmic vacuolation, and pericellular halo were the characteristic findings. Histopathology confirmed the diagnosis. A recent attempt to diagnose malignant CS on a recurrent lesion by FNAC was made in 2016 by Shobhanaa et al.³⁸ The cytology

	Hirsch and Helwig 1961 ²	Bekerecioglu et al 2002 ⁵	Yavuzer et al 2003 ⁴	Salama et al 2004 ¹³	Ayala-Cortes et al 2015 ³
Period	Not mentioned	1995–2001	1986–2002	1985–1977	1997–2014
No. of cases	188	13	16	25	19
Male/females	145/40	5/8	10/6	14/11	14/5
(M:F ratio)	(3.5:1)	(1:1.6)	(1.6:1)	(1.2:1)	(2.8:1)
	Unknown: 3				
Mean age	Not mentioned	33.1 years	42.8 years	55 years	50 years
(range)		(19-53 years)	(23–65 years)	(35–88 years)	(16.7 SD)
Mean size	Not mentioned	2.01 cm	Not mentioned	0.5 cm	0.9 cm
(range)		(0.8–3.1 cm)		(0.3–0.9 cm)	(0.47 IQR)
Sites					
Head and neck	150	10	15	16	17
Axilla and chest	6	0	0		0
Trunk	8	_	0	Others: 9	2
Extremities	19	2	_		0
Genitalia	2	0	0		0
No. of lesions	Single (except one case)	Single	Single	Single	Single
Most common clinical diagnosis	Sebaceous cyst or cyst	Not mentioned	Dermal cyst	Not mentioned	Cystic lesions or adnexal tumors
FNAC done	No	No	No	No	No
Treatment	Excisional biopsy	Excisional biopsy	Excisional biopsy	Excisional biopsy	Excisional biopsy
Abbreviations: FNAC, fine needle aspi	ration cytology.				

Table 1 Comparison of the clinical features of CS lesions in five large studies

	Age (years)/ gender	Site	Size (cm)	Duration (years)	Clinical diagnosis	Cytological diagnosis	Histopathological diagnosis
Masood	76/F	Left thigh	5×4	5	NM	CS	CS
et al 1988 ¹⁷							
Srinivasan	60/M	Right	2.5	3 months	Neurofibroma	CS	CS
et al 1993 ²²		shoulder					
Gottschalk-	82/F	Axilla	0.5×3	NM	Metastatic	Probable CS	CS
Sabag et al 1994 ¹⁹					lymph node		
Kumar et al 2003 ²¹	32/M	Nape of the neck	5×5×3	2	Hamartoma	Benign appendageal tumor of the skin	CS
Siddaraju	43/F	Dorsum of	0.8×0.8	1	Basal cell	CS	CS
et al 2009 ¹⁴		the nose			carcinoma		
Kumar	20/M	Dorsum of	2×2	NM	Dermoid cyst	CS	CS
201015		nose					
Skoro et al 2010 ¹⁶	63/M	Neck	0.8	5	NM	CS	CS
Dubb et al	32/F	Scalp	2	NM	NM	Suggestive of CS	CS
2010 ²⁰	23/M	Scalp	2				
	18/F	Upper lip	0.5				
Tokyol et al 2010 ²⁸	57/F	Philtrum	1.5	10	Lipoma	Benign appendageal tumor	CS
Nasit et al 2012 ²⁷	40/F	Mastoid	I.2 cm	3	None	CS	CS
Narasimha et al 2013 ¹²	50/M	Lower back	12×8×5	3	NM	CS	CS
Khan 2013 ¹⁸	31/M	Left	3×2.5	NM	Sebaceous or	Benign cystic neo-	CS
		supraorbital			epidermal cyst	plasm possibly	
		region				benign skin adnexal	
		_				tumor	
Pal et al 2014 ²⁹	33/M	Left forearm	2×1.5	1 1/2	NM	CS	CS
Barman et al 2016 ³⁹	25/M	Right thumb	3.5	2	NM	NM	CS
Rogers et al	67/M	Right axilla	1	1	Lymph node or	Benign epithelial-	CS
2016 ²⁴					cyst	mesenchymal bipha-	
						sic neoplasm	
Mahantappa et al 2016 ²³	40/M	Anterior abdominal	8×6×5	1/2	Dermoid cyst	CS	CS
		wall					
Lamba et al	37/M	Left arm	2.5×2	1	Epidermal inclu-	CS	CS
2017 ²⁶					sion cyst		
Our case	44/M	Upper lip	1.5×1.5	6	Sebaceous cyst	CS	CS
2018					and dermatofi-		
					broma		

 Table 2 Clinical findings and original diagnoses of cases which underwent FNAC

Abbreviations: CS, chondroid syringoma; FNAC, fine needle aspiration cytology; NM, Not mentioned.

Masood et al 1988 ¹⁷ NM Moderate Clusters and shees Small cells with relatively scant, fainty essionphi- lic cytoplasm Small ovoid-to- finely granular chromatin, occasional small chromo- centers NM Chondroid Srinivasan et al 1993 ²² NM NM NM NM MM MM MM MM Monomorphic configuration Some spindle- shaped cells Some spindle- shaped cells Myxoid Gottschalk- gadatious NM NM NM Single, groups and tubular configuration Regular NM NM Myxoid Siddaraju et al 2009 ¹⁴ Thick, mucoid and gelatinous NM Moderate Clusters as well as dispersed Round and monomorphous as dispersed Monomorphic erate to abundant cyto- plasm In clusters Metachrom, conspicuous Kumar 2010 ¹⁵ Mucoid NM Clusters an palicit to cor abundant cyto- plasm NM MM Relatively pr stained, cyto- plasm Kumar 2010 ¹⁵ Mucoid NM Clusters and palilary formations Round with moderate to abundant cyto- plasm NM MM Chondromy Skoro et al 2010 ¹⁶ Bloody NM Clusters and paplilary format		Aspirate	Cellularity	Epithelial cell arrangement	Individual cells	Nuclei	Myoepitheliod cells	Background/ Stroma
Srinivasan et al 1993 ²² NM NM NM Round to oval a moderate a moderate a mount of cytoplasm Monomorphic nuclei Some spindle- shaped cells Myxoid (Abundan) and path Gottschalk- Sbaag et al 1994 ¹⁹ NM NM Single, groups and tubular configuration Regular NM NM Myxoid Kumar et al 2003 ²¹ Thick, muccid and gelatinous Thick, muccid and gelatinous NM Moderate epithelial Round and monomorphous with morphores abundant amount of cytoplasm Monomorphic, much moderate to abundant amount of cytoplasm In clusters Metachroma chondromys Siddaraju et al 2009 ¹⁴ NM Moderate Round to poly- plasm NM NM Relatively pr stainee, cyan blink cronspicuous NM Relatively pr stainee, cyan philic to eosi philic groum plasm Some with moderate ani- abundant cyto- plasm NM Relatively pr stainee, cyan philic to eosi philic groum plasm Kumar 2010 ¹⁵ Mucoid NM Clusters Round with moderate to abundant crospicuous NM Chondromy corplasm Skoro et al 2010 ¹⁶ Bloody NM Clusters and papillary formations Well defined with dense, moderate cytoplasm NM Man Chondromy corplasm	Masood et al 1988 ¹⁷	NM	Moderate	Clusters and sheets	Small cells with relatively scant, faintly eosinophi- lic cytoplasm	Small ovoid-to- elongated with finely granular chromatin, occasional small chromo- centers	NM	Chondroid
Gottschalk- Sabag et al 1994 ¹⁹ NMNMSingle, groups and tubular configurationRegularNMNMMMMyxoidKumar et al 2003 ²¹ Thick, mucoid and gelatinousThick, mucoid and gelatinousThick, mucoid and gelatinousModerateRound and monomorphous with moderate to abundant amount of cytoplasmMonomorphic, main Some eccentrically of cytoplasmIn clustersMetachromy chondromy2Siddaraju et al 2009 ¹⁴ NMModerateClusters as well a slippersedRound to poly- gonal with mod- erate to abundant cyto- plasmNMRelatively pr stained, cyar plasmNMRelatively pr stained, cyar plasmKumar 2010 ¹⁵ MucoidNMClustersRound with moderate to abundant cyto- plasmConspicuous corplasmNMRelatively pr stained, cyar philic to eos abundant cyto- plasmNMRelatively pr sonucleosisKumar 2010 ¹⁵ MucoidNMClustersRound with moderate to abundant cellsCocasional rounded nuclei with clear halosNMChondromy chondromyKumar 2010 ¹⁵ BloodyNMClusters and papillary formationsVeil defined with dense, moderate cytoplasmNMChondromy chondromySkoro et al 2010 ¹⁶ BloodyNMClusters and papillary formationsVeil defined with dense, moderate cytoplasmNMChondromy chondromySkoro et al 2010 ¹⁶ BloodyNMClusters and papillary	Srinivasan et al 1993 ²²	NM	NM	NM	Round to oval cells with a moderate amount of cytoplasm	Monomorphic nuclei	Some spindle- shaped cells	Myxoid (Abundant)
Kumar et al 2003 ²¹ Thick, mucoid and gelatinousClusters of epithelialRound and monomorphousMonomorphic, with moderate to abundant amount of cytoplasmIn clustersMetachroma chondromysSiddaraju et al 2009 ¹⁴ NMModerateClusters as well as dispersedRound to poly- gonal with mod- erate to abundant cyto- plasmNMRelatively pr stained, cyar moderate ani- abundant cyto- plasmNMRelatively pr stained, cyar moderate or abundant cellsNMRelatively pr stained, cyar moderate or abundant cellsNMChondromy stained, cyar plasmKumar 2010 ¹⁵ MucoidNMClustersRound with moderate to abundant cytoplasmNMChondromy corestail papillary formationsNMChondromy cytoplasmSkoro et al 2010 ¹⁶ BloodyNMClusters and papillary formationsWell defined with dense, moderate cytoplasmNMChondromy cytoplasmSkoro et al 2010 ¹⁶ BloodyNMClusters and papillary formationsWell defined with dense, m	Gottschalk- Sabag et al 1994 ¹⁹	NM	NM	Single, groups and tubular configuration	Regular	NM	NM	Myxoid
Siddaraju et al 200914NMModerateClusters as well as dispersedRound to poly- gonal with mod- erate to abundant cyto- plasmOval, vesicular with mild to moderate ani- abundant cyto- plasmNMRelatively pr stained, cyar philic to eos 	Kumar et al 2003 ²¹	Thick, mucoid and gelatinous		Clusters of epithelial	Round and monomorphous with moderate to abundant amount of cytoplasm	Monomorphic, with fine chro- matin Some eccentrically placed	In clusters	Metachromatic, chondromyxoid
Kumar Mucoid NM Clusters Round with Monomorphic, NM Chondromy 2010 ¹⁵ August and the second seco	Siddaraju et al 2009 ¹⁴	NM	Moderate	Clusters as well as dispersed	Round to poly- gonal with mod- erate to abundant cyto- plasm A few occasional, tiny clusters of bland spindle cells	Oval, vesicular with mild to moderate ani- sonucleosis Some with conspicuous nucleoli and chromocenters Occasional rounded nuclei with clear halos	NM	Relatively pale- stained, cyano- philic to eosino- philic ground substance
Skoro et al 2010 ¹⁶ Bloody NM Clusters and papillary Well defined with dense, moderate formations Round to NM Chondromy Image: Clusters and papillary Clusters and dense, moderate formations ovale, centrally Image: Clusters and ovale, centrally Image: Clusters and ovale, centrally Image: Clusters and ovale, centrally Image: Clusters and ovale, centrally Chondromy	Kumar 2010 ¹⁵	Mucoid	NM	Clusters	Round with moderate to abundant cytoplasm	Monomorphic, centrally to eccentrically located Fine chromatin	NM	Chondromyxoid
chromatin	Skoro et al 2010 ¹⁶	Bloody	NM	Clusters and papillary formations	Well defined with dense, moderate cytoplasm	Round to ovale, centrally located Fine, evenly distributed chromatin	NM	Chondromyxoid

Table 3 Detail cytological findings of cases which underwent FNAC

	Aspirate	Cellularity	Epithelial cell arrangement	Individual cells	Nuclei	Myoepitheliod cells	Background/ Stroma
Dubb et al 2010 ²⁰	NM	NM	Sheets, clusters and single cells	Well defined with moderate eosi- nophilic to amphophilic cytoplasm imparting a plasmacytoid appearance	Bland, round to oval, eccentri- cally located Evenly dis- persed, hypo- chromatic chromatin Small, incon- spicuous nucleoli.	NM	Eosinophilic myxoid
Tokyol et al 2010 ²⁸	NM	Hypercellular	Cohesive groups of cells	Monomorphic round cells with moderate to abundant amount cytoplasm	Monomorphic nuclei with fine chromatin. Some nuclei were eccentri- cally placed, like plasmacy- toid cells	Spindle cells seen	Chondromyxoid
Nasit et al 2012 ²⁷	Thick and mucoid	NM	Sheets and loose clusters with a few single cells	Bland, small and monomorphic with moderate amount of cytoplasm	Round to-oval, centrally located Evenly dispersed fine chromatin	Elongated	Chondromyxoid
Narasimha et al 2013 ¹²	Thick, mucoid, and gelatinous	NM	Loose cohesive clusters and discretes	Round to oval with moderate to abundant cytoplasm	Centrally located nuclei having fine chromatin, a few showing one to two prominent nucleoli	NM	Chondromyxoid
Khan 2013 ¹⁸	Thin fluid- like	Moderate	Cohesive clus- ters A few acinar formation	Medium-sized cells with moder- ate to abundant amount of cytoplasm	Bland appearing monomorphic centrally placed or slightly eccentric nuclei with fine chromatin	Smaller hyperchromatic	Chondromyxoid (Scant)
Pal et al 2014 ²⁹	Thick mucoid	Moderate	Clusters	Monomorphic, round to oval, medium sized having moderate amount of cytoplasm	Bland round to oval with finely dispersed chromatin	Small cells having plasmacytoid appearance with dark nuclei	Chondromyxoid (Abundant)

(Continued)

, Table 3 (Continued).

	Aspirate	Cellularity	Epithelial cell arrangement	Individual cells	Nuclei	Myoepitheliod cells	Background/ Stroma
Barman et al 2016 ³⁹	NM	NM	Loose clusters and sheets	Ovoid and spin- dle Cells with moderate to the abundant well defined cytoplasm	Oval with bland finely granular chromatin	NM	Chondromyxoid
Rogers et al 2016 ²⁴	NM	Moderate	Loose or clusters	Epithelioid to spindled with a moderate amount of cytoplasm	Round to ovoid nuclei, and inconspicuous nucleoli		Myxoid
Mahantappa et al 2016 ²³	Scant and Mucoid	Scant	Clusters, groups, in papillae	Small to medium size with well- defined cell bor- ders having scant- to-moderate amounts of cytoplasm	Round to oval with fine stippled chromatin	Clusters	Myxochondroid
Lamba et al 2017 ²⁶	Thick mucoid	NM	NM	Monomorphic, round to oval, with moderate amount of cytoplasm	Centrally placed nuclei with fine chromatin	NM	Chondromyxoid
Our case 2018	Blood mixed		Aggregates, acini and singly scattered	Round to poly- gonal with baso- philic dense moderate cytoplasm	Central to eccentric, round to oval nuclei with bland chromatin	Aggregates	Chondromyxoid

Abbreviations: NM, not mentioned.

revealed hypercellularity and tissue fragments of malignantappearing round-to-polygonal cells. The biopsy was inconclusive. A repeat FNAC was performed, along with immunocytochemistry. Vacuolation, indistinct cell borders, nuclear pleomorphism and multiple prominent nucleoli were appreciated. Pan cytokeratin, EMA, S-100, calponin, and α -smooth muscle actin showed strong positivity, which sealed the diagnosis.

Excision is the treatment of choice and should include the margins. In 1961, Hirsch and Helwig² proposed the histological criteria for diagnosis of CS. Apocrine CS exceeds the number of eccrine CS reported. The apocrine variant has two rows of epithelial cells lining the tubular and cystic branching lumina, while the smaller lumen in eccrine type has a single row of cells.⁴ The presented case belonged to the former group.

Conclusion

CS is a rare tumour presenting in dermatological practice. FNAC is a very useful tool for making preliminary diagnosis of CS before making a large excision. However, the final diagnosis is based on histopathological examination.

Ethical statement

The patient gave his written informed consent for the publication of images and information. Institutional approval was not required to publish the case details.

Disclosure

The authors report no conflicts of interest in this work.

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