

Research Article

Diagnosis of Mercurial Teeth in a Possible Case of Congenital Syphilis and Tuberculosis in a 19th Century Child Skeleton

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Without the presence of "*caries sicca*," "sabre shins," and nodes/expansion of the long bones with superficial cavitation, differential diagnosis of venereal syphilis and tuberculosis (TB) may be difficult as various infections produce similar responses. However, congenital syphilis has distinctive features facilitating a diagnosis. A case study of remains of a juvenile European settler (probably male, 8–10 years old) (B70) buried in the 19th century and excavated in 2000 from the cemetery of the Anglican Church of St. Marys in South Australia is presented. B70 demonstrated that the two diseases might have been present in the same individual, congenital syphilis and TB. Widespread destruction of vertebral bodies and kyphosis-related rib deformations indicate advanced TB. Severe dental hypoplasia is limited to permanent incisors and first molars; there is pitting on the palate, periosteal reaction on the skull vault, and thinned clavicles. Dental signs are not limited to "screwdriver" central incisors and mulberry molars. Apical portions of the crowns of permanent upper, lower, central, and lateral incisors have multiple hypoplastic-disorganized defects; deciduous canines have severely hypoplastic crowns while possibly hypoplastic occlusal surfaces of lower deciduous second molars are largely destroyed by extensive caries. These dental abnormalities resemble teeth affected by mercurial treatment in congenital syphilitic patients as described by Hutchinson.

1. Introduction

In the past, the presence of numerous diseases and the lack of an effective form of treatment meant individuals could have suffered from more than one disease. This is especially the case in relation to chronic afflictions that could be combined with congenital diseases or acute infections. Syphilis and tuberculosis (TB) were two of these diseases. Significant in the past, both diseases continue to be an important public health problem. Syphilis, caused by the spirochete Treponema *pallidum* is typically transmitted through sexual contact. It can also be transmitted via the placenta from an infected mother to the fetus while she is in the most infectious stages of the disease (early primary or secondary stage). It is known as congenital syphilis [1]. Syphilis affects more than 12 million adults [2-4] and a million pregnancies each year [4-6]. Tuberculosis, a chronic infectious disease caused by Mycobacterium tuberculosis, is usually transmitted through the inhalation of airborne droplets filled with bacteria produced by infected individuals usually when coughing [7, 8]. Approximately 9 million new cases were registered and 1.5 million people died from tuberculosis in 2013 [9].

In most palaeopathological studies, skeletal signs of diseases are diagnosed to one nosological unit. This finds some justification in the fact that only a small portion of diseases leave recognizable signs on hard tissues of the body (bones and teeth). It is, however, possible to find signs of more than one affliction on a single skeleton [10]. When this is the case, study of skeletal involvement should not be the only method applied when making a differential diagnosis.

The differential diagnosis of syphilis and tuberculosis in palaeopathological specimens remains difficult as both diseases rarely affect or leave any signs on hard tissues of the body. In syphilis, only 1/3 of individuals suffering from the tertiary stage of the disease will develop any bone lesions [10] while only about 3% to 5% of individuals with active TB will have skeletal changes [11–13]. The diagnostic characteristic of syphilis include "caries sicca," sclerosis, and pitting of the outer table of the cranial vault resulting from accumulation of stellate scarring [11, 14] creating a "worm eaten" appearance [14], tibial bowing, known as sabre shin [11, 15, 16], and the expansion of the long bones with nodes with superficial cavitation [14]. In tuberculosis diagnostic elements include osteolytic lesions on the thoracic and lumbar vertebral bodies [10, 13, 17]. Rib involvement including new bone formation, particularly periosteal reactions on the visceral surface [18– 21] is now considered in the diagnosis of tuberculosis [22].

Lesions of congenital syphilis can also be difficult to identify in skeletal samples as many pregnancies can result in stillbirths, abortion, or death [15, 23] and those skeletons are not often preserved. However, in those patients that do survive, the disease causes a disturbance in dental development producing abnormalities that are distinguishable features of the disease. The most recognisable are Hutchinson's incisors, while others include Moon's molars and Fournier's "mulberry" molars [15, 16, 24–30]. It is this characteristic that can support a differential diagnosis of the disease.

However, in cases where these diagnostic changes are not present, differential diagnosis of a specimen can be difficult. Our knowledge of the type of treatments used to combat syphilis and tuberculosis throughout history is well known. They used natural remedies, chemical compounds, and recently penicillin; however, our knowledge of the effects of these treatments on hard tissues has not been explored in depth.

Mercury has been used as early as the 27th century BC in China [31]. It was recognised as a form of treatment for venereal diseases [31-33] prior to the introduction of salvarsan [34–37] and penicillin in the 20th century [38]. Mercury was provided to mothers during pregnancy [39] children, and infants in the form of ointments, calomel teething powders [24, 40, 41], and injections [41, 42]. Mercurial poisoning was noted by Sir Hutchinson [24, 25, 40] to grossly influence tooth development producing abnormalities of enamel formation (Figure 1). These may interfere with the expression of "classic" dental signs of congenital syphilis. When salvarsan was introduced, replacing mercury early in the 20th century, American military physicians recommended the use of mercury for the treatment of tuberculosis in adult patients [43, 44], but it is unclear how widespread this method of TB treatment became. There is no mention of its effects on dentition.

This paper presents a case study of the pathological lesions observed on a European subadult dated from the mid-19th to early 20th centuries who died during the early European colonization of South Australia, Australia [45]. The influences of mercury are considered in this case. In order to understand variation in skeletal lesions it is useful to consider the treatments used and their possible effects on the hard tissues of the body. This method may assist in a differential diagnosis.

2. Materials and Methods

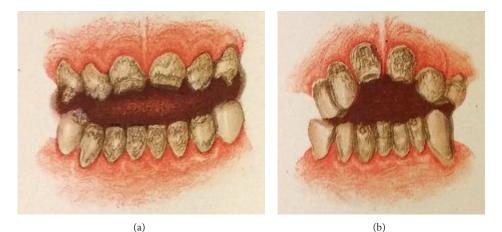
The juvenile in this study (B70) was among a sample of 70 individuals excavated in 2000 from the cemetery of the Anglican Church of St. Marys, located at 1167 South Road, in St. Marys, Adelaide, South Australia. Many of those buried at the cemetery were in unmarked graves in a section of the grounds dating from 1846 to 1927, preventing

individual identification [46]. These unmarked graves were considered colloquially as "paupers" graves due to their low socioeconomic status. Written records of burials can be found at the Church's Office. Signs of various infections were found on paleopathological analysis among the skeletal sample excavated including acquired syphilis, tuberculosis, pulmonary, and systemic infections. Some of these were also listed as causes of death in parish records [46]. Two thirds of the skeleton survives (Figure 2). Bone tissue is fragile and poorly preserved with some bones missing and others in fragments. The individual was aged by dental development, eruption, and formation using the Ubelaker chart [47] and primary ossification centres [47, 48]. Sex of a subadult is difficult to estimate [49-51] and the methods proposed do not produce highly reliable results. Using the morphology of the symphyseal region of the mandible [52] and the shape of the mandible [53] in combination with the robusticity of long bones [54], the shape of the sciatic notch would have been used to aid in determining sex; however, the majority of the pelvis is missing. To determine the effects of mercury on hard tissues and possible pathologies, a search of the literature was conducted and compared to B70.

3. Results

B70 is probably a subadult male. According to dental eruption and formation, the child is between eight and ten years of age. An osteoblastic lesion approximately 15 mm in diameter is present on the cranial vault (possible periosteal reaction) on the posterior portion of the left parietal bone close to the lambdoid suture (Figure 3). Pitting is present on the maxillary alveolar process (Figure 4) and on both sides of the palate; however, it is stemming from the root of the right first upper molar (Figure 5).

3.1. Dentition. B70 demonstrates mixed dentition. The maxillary upper right central incisor is the only tooth missing postmortem. The dentition consists of a left central incisor, partially erupted lateral right and left incisors, deciduous canines, first and second premolars, permanent first molars, and second permanent molar germs. The maxillary left central incisor demonstrates narrow and rounded medial and distal edges and is slightly crescentic in shape. It is hypoplastic. Its incisive edge is slightly narrowed with minute mamelons and multiple notching. The incisal 1/3 of the labial surface has thinner, discoloured (darker) enamel with pitting hypoplasia. This part of the crown forms a few months after birth [55]. The remainder of the labial surface has three transverse hypoplastic lines (Figure 4). On the lingual surface, the incisal 1/3 of the crown has thinner enamel. It is separated from the remainder of the crown by a distinct hypoplastic grove that extends to the mesial and distal surfaces. The lateral right incisor is narrow and hypoplastic. The right and left maxillary lateral incisors on the labial view, approximately a third of the distance from the apical point of the crown, are a round indentation in the enamel. The lateral left incisor has a central pit about 1mm in diameter and is notched mesially. Crowns of the upper deciduous canines have wide hypoplastic discoloured (darker) areas beginning below



(c) (d)

FIGURE 1: Diagrams of mercurial teeth seen in mercurial treated congenital syphilitic patients by Hutchinson. Hutchinson, J. 1878. Illustrations of clinical surgery consisting of plates, photographs, woodcuts, diagrams, and so forth: illustration of surgical diseases, symptoms and accidents, also operative and other methods of treatment, with descriptive letterpress, London, J. & A. Churchill.

the tip of the crown and extending down to about 1/3 of the crown indicating that the changes occurred after birth (Figures 4 and 5). All maxillary premolars appear normal. Both first permanent upper molars have grossly abnormal crowns. Their occlusal surfaces have widespread hypoplastic defects (Figures 6(a) and 6(b)). Extensive carious lesions are present on the mesial half of the occlusal surface of the right upper first permanent molar and small carious lesions on the occlusal surface of the left first permanent molar. Distinctive lines of thinner enamel are present on both permanent molars, separating the upper part of the crown (occlusal surface) from the rest of the crown. Areas constricted by the lines are smaller than the extent of

(e)

the lower parts of the crowns. This indicates that the changes occurred shortly after birth [55]. The crown morphology of the second permanent molar germs is normal.

(f)

Mandibular dentition includes all permanent incisors, deciduous molars, canines, first permanent molars, and second permanent germs. The permanent incisors are hypoplastic with small mamelons and linear and pitted hypoplasia on the crowns (Figure 7(a)). Distal 2/3 crowns of lower deciduous canines are very narrow with thinner hypoplastic enamel and appear conical in shape (Figure 7(a)). The remaining proximal 1/3 has rather normal enamel. The proximal 1/3 of deciduous canine crowns has normal enamel. First deciduous molars show discolouration but no caries.

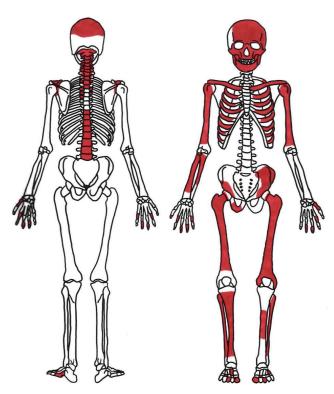


FIGURE 2: Shaded areas represent bones present.



FIGURE 3: Periosteal reaction approximately 15 mm in diameter on posterior portion of the left parietal bone close to the lambdoid suture. On close inspection there is no erosion of the cortical bone (Lamina externa).

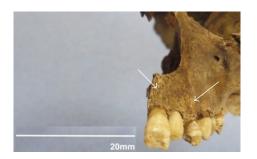


FIGURE 4: Pitting on maxillary alveolar process due to inflammatory response.



FIGURE 5: Pitting on palate stemming from permanent first molar due to inflammatory response.

The second deciduous molars and canines have extensive carious cavities. The first permanent molars occlusal surfaces are grossly hypoplastic (Figure 7(b)). Similarly to the maxillary dentition, hypoplastic changes indicate that they have occurred within the first few months after birth. The right lower permanent molar has an extensive carious lesion extending through most of the centre of the occlusal surface. A small carious pit in the centre of the mesial half of the occlusal surface is present in left (Figure 7(b)). The crown morphology of both lower second molar germs is normal. Resorption of the alveolar bone is observed (Figure 7(a)).

3.2. Clavicle and Ribs. Morphology of the clavicle and several ribs appears abnormal. Thinning of the sternal end of the clavicle is evident (Figure 8). There is a small proliferative change on the upper portion of the 3rd rib. Localised inflammatory reaction is present on the right side, superior surface on the 4th or 5th rib (Figure 9). Added grooving is evident on the superior surface of several ribs.

3.3. Vertebral Column. There are extensive pathological changes on the vertebral column. The vertebral bodies of C5-Th3 show damage to their anterior parts. Signs of remodelling on C6 and C7 could indicate signs of healing (Figure 10). Cervical vertebrae C1-C4 show no pathological signs. Vertebral bodies of Th3-Th4 are largely destroyed, Th4 more so than Th3. Bodies of all other thoracic vertebrae, except Th10 and Th11, are absent, but it cannot be ascertained whether this was due to taphonomic processes or due to actual pathological destruction. Zygapophyseal joints between what are likely to be Th5-Th6 are completely fused on both sides and there are no vertebral bodies (Figure 11(a)). The left zygapophyseal joints of Th6-Th7 are also fused, while the right side is missing. Th9 is possibly in fragments. Vertebral body of Th10 is partially destroyed. Th11 and Th12 are represented by small fragments.

Two bodies and two arches of the lumbar vertebrae are preserved. One body has two deep pits on its anterior surface, which appear lytic (Figure 11(b)). The other body and the arches show no pathological signs. The body and right lateral mass of the first sacral segment are preserved without obvious pathological signs. The left lateral mass of the sacrum

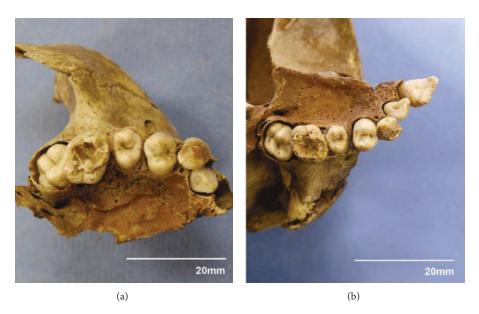


FIGURE 6: (a) Right permanent upper first molar showing signs of dysplastic occlusal enamel. (b) Left permanent upper first molar with hypoplastic defects characteristic of dysplastic occlusal enamel.



FIGURE 7: (a) Angled lower border of mandible, similar in shape to male juvenile mandible A.668 studied by Loth and Henneberg [52, Figure 2]. (b) Lower first permanent molars grossly hypoplastic.



FIGURE 8: Thinning at sternal ends of clavicles.

is completely fused with the left ilium at the sacroiliac joint. There are no clear signs of any inflammatory processes. Right sacroiliac joint appears normal. The first sacral segment has a normal body. Bodies of other sacral segments are preserved in fragments and no pathological signs were observed. No pathological signs were noted on the long bones.



FIGURE 9: Inflammatory reaction of inferior surface on 4th or 5th rib.

4. Discussion

4.1. Differential Diagnosis. In this case, it is possible that B70, a mid-19th to early 20th century specimen, suffered from multiple conditions. Differential diagnosis of B70 includes



FIGURE 10: Destruction of anterior bodies of lower cervical vertebrae.



(a)



FIGURE 11: (a) Fusion of zygapophyseal joints between Th5 and Th6 completely fused with no vertebral bodies. (b) Lumbar vertebral body with deep pits.

infectious and noninfectious diseases including congenital syphilis, tuberculosis, brucellosis, rickets, and fluorosis.

Lesions in congenital syphilis can vary from periosteal reactions and osteomyelitis in the early stages [56–58] and cranial gummatous lesions and frontal bossing of the bone, destruction of the nasal bridge, a high arch palate, sternoclavicular thickening, and tibial bowing (sabre shin) in the late stage of the disease [15, 16, 59–61]. With the exception of a possible localised periosteal reaction on the cranial vault of B70, there is minimal skeletal evidence to support the differential diagnosis of congenital syphilis.

The dental changes in B70, although not "typical" (Hutchinson's incisors, Moon's molars, or Fournier's mulberry molars), may still be a result of congenital syphilis

through the mercurial treatment of the disease. Hutchinson recognised that mercury produced enamel defects in particular pairs of teeth. In severe cases it would affect dentine, too. With the tooth enamel deficient, the tooth would appear rugged, pitted, and dirty [40]. The first permanent upper and lower molars are the "test teeth" for mercurial influence, similar to the upper central incisors considered to be the "test teeth" in congenital syphilis. The crown enamel is deficient, with dentin growing through and revealing numerous discoloured tubercles [40]. A distinct demarcated line separating healthy enamel from diseased enamel was also evident on the sides of the molars. In severe cases, the tooth could appear dwarfed. Upper and lower incisors and canines were usually affected, with enamel deficiencies occurring below a line that would cross them at the same level; however, premolars usually escape all damage [40]. Hutchinson also noted that it was common for both syphilitic and mercurial teeth to be present at the same time which may have caused confusion among physicians [40]. He did not specify, however, exact ages or developmental stages of dentition at which the changes occurred.

B70's dentition closely resembles the descriptions and images (Figure 1(c)) of congenital syphilis patients treated with mercury as provided by Hutchinson. The first upper and lower permanent molars demonstrate enamel deficiency across the occlusal surface exposing multiple tubercles, appearing rugged, pitted, and dirty. There is a clear distinction between diseased and healthy enamel on all four molars and all three canines. All upper and lower incisors exhibit enamel deficiencies apical to the linear enamel hypoplasia. All upper premolars appear normal. Taking into account individual variation in formation of deciduous and permanent tooth crowns, the most likely age at which the changes in B70's dentition occurred is shortly after birth. The cervical ends of enamel on all teeth appear normal, suggesting that ameloblasts were disturbed during the early years of life [62]. Tips of deciduous canine crowns seem to be normally formed, but the crown area below them is hypoplastic in contrast to first deciduous lower molars whose morphology is normal. Permanent tooth changes affected apical or occlusal portions of specific crowns that form in the first few months of life. It is possible that the type of enamel damage to the first permanent molars in B70 could be classified as cuspal enamel hypoplasia [62]; however, to confirm this, scanning electron microscopy would need to be performed.

Clinical presentations of congenital syphilis present similar dental features to those seen in B70. These include multiple notching or serrated edges which were seen in five patients [63] pitted enamel hypoplasia of the upper central and lateral incisors and primary and secondary dental caries on numerous teeth [64]. Narrowing and a reduction of the dentinoenamel junction of the permanent incisors and first molars, with a reduction in the size of the crowns and constriction of mamelons, was also noted by Sarnat and Shaw [27].

In comparison to palaeopathological specimens, similarities to B70 include the round indentation in the enamel on the maxillary right and left lateral incisors [65–67] and pitted enamel hypoplasia in the lower right incisors. Others include linear enamel hypoplasia in all four incisors with a deficiency in enamel above (apically) a hypoplastic line [67, 68], the distinct demarcation between healthy and diseased enamel, and severe enamel deficiencies exposing multiple tubercles in molars [67].

The lack of skeletal lesions on limb bones of B70 could be supported by clinical cases of late congenital syphilis, which found no periosteal lesions or perichondritis [69, 70]. This may be related to the stage of infection in maternal syphilis and transmission [69]. The later stages of the disease in the mother produce lesser risk of infection [15] and possibly less severity.

Tuberculosis is typically diagnosed by osteolytic skeletal lesions in the vertebral bodies and in large joints of palaeopathological specimens [11]. The most common manifestations of skeletal tuberculosis in children are spondylitis, osteomyelitis, and involvement of the joints [11, 71]. In children, common areas affected by the disease included the knee [72, 73], lytic circumscribed lesions of the cranium [7, 74, 75], spine [71, 76], hip [73, 77], elbow [78], and ribs [76]. There is no documentation with regard to dental abnormalities found in juvenile tuberculosis [7, 79]. Dental changes briefly mentioned include linear enamel hypoplasia [80, 81], carious lesions, and decreased enamel thickness [80].

Comparing B70 to skeletal signs of tuberculosis, osteolytic lesions evident on the thoracic and lumbar vertebrae resemble few juvenile specimens [81, 82]. A circumscribed periosteal lesion on the superior surface of rib four or five in B70 is similar to that found in the case of TB in the Hamann-Todd Osteological Collection [83]. However, no lytic lesions were apparent on the cranial vault of B70, neither was there involvement of the joints as in the cases mentioned above. Linear enamel hypoplasia and dental abnormalities as seen in B70 have not been noted in clinical cases of primary tuberculosis [84-86]. There are no documented palaeopathological cases of congenital tuberculosis. This may be due to the rarity of the disease and the low survival rates of infants born with the condition [87-89]. Therefore, congenital TB is not known to produce extensive hypoplastic defects on incisal edges nor on occlusal surfaces of teeth. It is likely that B70 suffered from TB acquired during childhood.

While we know that mercury has been used in the treatment of tuberculosis, its descriptions and suggested use begin from 1908 and they do not seem to be widespread. B70 was buried in a cemetery dating from 1846 to 1927, so it is unlikely that mercury's use in treatment of tuberculosis would be the cause of described dental changes.

Brucellosis affects different areas of the skeleton in adults and in children. In adults the spine or sacroiliac joint is more commonly affected, whereas in children, the knee, hip, and ankle joints are more common [90–94]. While the left sacroiliac joint is fused in B70, who is a child, there do not appear to be any signs of inflammation and there are no other pathologies that resemble those seen in brucellosis; therefore, it is difficult to make a confident differential diagnosis. However, the sacral segments that are present do not show any pathology. There are also no lesions present on the knee joint or the rest of the appendicular skeleton, and thus brucellosis is unlikely. Rickets is a vitamin D deficiency, affecting the metabolism of calcium and phosphorus and the mineralization of bone. Skeletal changes include bending deformities [95, 96], metaphyseal flaring, and porosity of cortical bone [11, 95, 97]. These changes can affect the cranial vault, long bones, pelvis, ribs, and vertebrae. In conjunction with the skeletal pathologies of rickets, abnormalities in dentition are common, particularly linear enamel hypoplasia, pitting, dental opacities, and caries [98–100]. Considering that there are no bending deformities, flaring, porosity of the cortical bone, and dental opacities and while hypoplasia is not limited to linear defects, rickets in B70 is unlikely.

Fluorosis is a disturbance of dental development resulting from ingestion of large quantities of fluoride [11, 101]. These dental abnormalities include opaque white patches in the enamel. This can result in pitting, striations, and widespread brown staining [101–104]. Skeletal pathologies include abnormal bone formations on the appendicular or axial skeleton, mostly linked with the insertions of tendons and ligaments [11]. In clinically diagnosed cases of fluorosis in children, skeletal manifestations included osteopenia, growth lines, and sclerosis [105–107]. Considering that there is no widespread dental staining nor skeletal lesions relating to fluorosis, it is unlikely that B70 suffered from fluorosis.

5. Conclusion

B70 was excavated from St. Marys cemetery, from a section of the grounds dating from 1846 to 1927, when European settlers colonized South Australia. B70 was buried at the expense of the Government in a section of the cemetery referred to as the "paupers" graveyard [108]. Burial records of St. Marys indicate that treponemal diseases and tuberculosis were present among the skeletal sample B70 originated from and other skeletons (B10, B6, and B53c), demonstrated possible cases of treponemal disease [46]. Considering that B70 was excavated from the pauper's section of the graveyard and multiple diseases were present in the sample (syphilis and tuberculosis), it is probable that B70 suffered from multiple diseases in congenital syphilis and tuberculosis. The significance of this skeleton is that it displays dental signs that are not typically seen in congenital syphilitic cases. It is possible that this specimen displays the effects of mercury that was used to treat the disease. It is possible that chemical elements or compounds have not been considered in paleopathology to have an effect on hard tissues. Hopefully, this paper will reintroduce an interest in the work of Hutchinson who noted that mercury, used to treat syphilis, plays a role in the disruption of enamel formation. Mercury's effects are separate from tooth development (size and shape), caused by the disease and yet they are indicative of the disease through its treatment. Therefore, Hutchinson's incisors, Moon's molars, and Fournier's molars are not the only dental abnormalities that should be considered in the diagnosis of syphilis when examining specimens from antiquity up to the introduction and usage of modern treatments.

Conflict of Interests

The authors declare that there is no conflict of interests regarding the publication of this paper.

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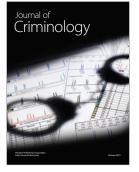
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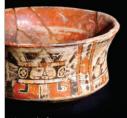






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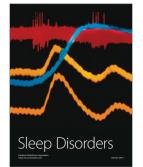




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