

Luigi Matturri · Giulia Ottaviani · Anna M. Lavezzi

Unexpected sudden death related to medullary brain lesions

Received: 10 February 2005 / Revised: 10 February 2005 / Accepted: 11 February 2005 / Published online: 18 May 2005
© Springer-Verlag 2005

Lesions of the tractus solitarius nucleus, generally of a congenital nature, may be the pathophysiological substrate of the sudden infant death syndrome (SIDS). The neuropathological studies we have conducted on a large population of victims of SIDS, of unexpected neonatal deaths and fetal deaths at term, revealed analogous congenital lesions of the structures of the brainstem that modulate respiratory activity, as well as cardiovascular functions, arousal and upper digestive activities [1, 2, 3].

The common denominator of all these deaths was the absence of neurological symptoms, generally associated with the presence of congenital lesions and in some cases of acquired lesions. In our studies, we have observed a wide variability of degree and localization of congenital lesions, such as hypoplasia/agenesis of the arcuate nucleus (present in about 50% of the cases we examined), associated with hypoplasia of the reticular formation in fetuses, hypoplasia of the parabrachial/Kölliker-Fuse complex, lesions of the nucleus of the solitary tract, gliosis (an expression of acute lesions or chronic injury), as well as functional alterations of the neurotransmitters, such as catecholamines in the locus coeruleus and somatostatin in the hypoglossus nucleus [3, 4, 5, 7, 8, 9].

We are in agreement with Jaster et al. [10] that subtle lesions involving the solitary tract nucleus, and other nuclei (particularly the arcuate nucleus and the Kölliker-Fuse nucleus), are responsible for disruption of neural pathways, and consequently for sudden perinatal death (of both the term fetus and the newborn) and early infant death without any premonitory neurological dysfunctions.

Our findings of a wide range of acquired lesions, also involving the solitary tract nucleus, determining unexpected infant death, raise very interesting issues. In particular, we have observed a T lymphocytic leptomeningitis affecting the ventral medullary surface [7], two cases with encephalitic features (probably of viral etiology) [9], and a hemangioendothelioma identified in the area postrema, which widely infiltrated the postero-middle brainstem bilaterally, sparing only the dorsal vagus nucleus [8]. Three cases of sudden adult death due to prions were also described [11].

As the volume of data on new morphological and functional alterations of the cardiorespiratory centers of the brainstem accumulates, it becomes ever more clear that it is essential that sudden death victims be submitted to an in-depth necropsy examination, focusing particularly on the brainstem in serial sections [1, 3, 8].

At the 7th International Conference on SIDS in 2002, we proposed [6] that the current definition of SIDS as “the sudden unexpected death of an infant under 1 year of age which remains unexplained after a thorough case investigation, including the performance of a complete autopsy, examination of the death scene, and review of the clinical history” can be effective only if the complete autopsy includes an in-depth histological examination of the cardiorespiratory innervation and specialized myocardium, performed by an experienced pathologist, as described on our web site: http://users.unimi.it/~pathol/sids_e.html.

References

1. Lavezzi AM, Ottaviani G, Ballabio G, Rossi L, Matturri L (2004) Preliminary study on the cytoarchitecture of the human parabrachial/Kölliker-Fuse complex, with reference to sudden infant death syndrome and sudden intrauterine unexplained death. *Pediatr Dev Pathol* 7:171–179
2. Lavezzi AM, Ottaviani G, Matturri L (2004) Role of somatostatin and apoptosis in breathing control in sudden perinatal and infant unexplained death. *Clin Neuropathol* 23:304–310

L. Matturri (✉) · G. Ottaviani · A. M. Lavezzi
“Lino Rossi” Research Center for the study and prevention of unexpected perinatal death and SIDS, Institute of Pathology, University of Milan, Via della Commenda 19, 20122 Milan, Italy
E-mail: luigi.matturri@unimi.it
Tel.: +39-02-5419521
Fax: +39-02-54198538

3. Lavezzi AM, Ottaviani G, Mauri M, Maturri L (2004) Hypoplasia of the arcuate nucleus and maternal smoking during pregnancy in sudden unexplained perinatal and infant death. *Neuropathology* 24:284–289
4. Lavezzi AM, Ottaviani G, Mingrone R, Maturri L (2005) Effects of smoking on human locus coeruleus development analyzed in perinatal and sudden unexplained death. *Dev Brain Res* 154:71–80
5. Maturri L, Lavezzi AM, Cappellini A, Ottaviani G, Minoli I, Rubino B, Rossi L (2003) Association between pulmonary hypoplasia and hypoplasia of arcuate nucleus in stillbirth. *J Perinatol* 23:328–332
6. Maturri L, Lavezzi AM, Rossi L (2002) Proposal to modify the definition of SIDS, with regard to the post mortem exam. In: *Proceedings of the 7th SIDS International Conference*. Florence, Italy, p 103
7. Maturri L, Ottaviani G, Ramos SG, Biondo B, Rossi L (1998) Discrete T-lymphocytic leptomeningitis of the ventral medullary surface in a case of sudden unexpected infant death. *Adv Clin Pathol* 2:313–316
8. Maturri L, Ottaviani G, Rossi L (1999) Sudden and unexpected infant death due to an hemangioendothelioma located in the medulla oblongata. *Adv Clin Pathol* 3:29–33
9. Ottaviani G, Maturri L, Bruni B, Lavezzi AM (2005) Sudden infant death syndrome “gray zone” disclosed only by a study of the brain stem on serial sections. *J Perinat Med* 33:165–169
10. Jaster JH, Zamecnik J, Bartos A, Dohan FC Jr, Smith TW (2005) Unexpected sudden death caused by medullary brain lesions involves all age groups and may include ‘sudden infant death syndrome’ as a subset. *Acta Neuropathol* (this issue) DOI 10.1007/s00401-005-0996-6
11. Valli G, Rossi L, Varesi C, Maturri L (1994) Brain stem involvement and long QT interval transmissible spongiform encephalopathy. *Ann N Y Acad Sci* 724:363–366