

KANET in Brazil: First Experience

Raul Moreira Neto

ABSTRACT

Objective: The goal of this pilot test is to compare Kurjak's antenatal neurological test (KANET) results between high and low-risk pregnancies and see the feasibility of using this test on a larger scale in Brazil.

Methods: We applied KANET test in high-risk ($n = 17$) and low-risk ($n = 34$) pregnancies and compared the results.

Results: There was a significant difference between groups of high-risk and low-risk pregnancies for all parameters in KANET score 2. For KANET score 0, 5 out of 8 parameters were significant different: isolated head anteflexion, cranial sutures and head circumference, isolated hand movement or hand to face movements, isolated leg movement and fingers movements. All abnormal KANET result came from high-risk pregnancies (17.6%). No low-risk pregnancies presented KANET score 0. Efficacy and effectiveness were not evaluated due to the imprecision inherent in data from small samples.

Conclusion: This successful preliminary study of KANET in Brazil showed important differences in fetal behavior between the low and high-risk pregnancies, and can be used as information of feasibility and identify modifications needed in the design of a larger testing study.

Keywords: Fetal behavior, Four-dimensional ultrasound, KANET, Neurodevelopment prenatal assessment, Fetal central nervous system.

How to cite this article: Neto RM. KANET in Brazil: First Experience. *Donald School J Ultrasound Obstet Gynecol* 2015;9(1):1-5.

Source of support: Nil

Conflict of interest: None declared.

INTRODUCTION

Over the past 30 years, emerged the most important results on fetal behavior, through the evolution of ultrasound and important research of basic neurological development. It is increasingly evident that many severe neurological disorders, or minimal cerebral dysfunctions, originate from the intrauterine, instead of the perinatal or early postnatal period.^{1,2}

After fetal behavior has been compared with morphological studies, investigators have come to conclusion that

fetal behavioral patterns directly reflect developmental and maturational processes of fetal central nervous system (CNS). Since, fetal brain impairment could interfere with the internal motor activity and fetal movements (or behavior) this could be used as a marker for fetal brain optimality.³⁻⁷ Therefore, it was proposed that the evaluation of fetal behavior could give the possibility to discriminate between normal and abnormal brain development, likewise an early diagnosis of different structural or functional abnormalities.³

The challenge of modern perinatal medicine is to assess the integrity of fetal and neonatal CNS especially since many important functional neurological abnormalities such as cerebral palsy (CP) are still poorly understood.⁸

The discovery of the continuity of neurological activity generated from the prenatal period to postnatal life, allowed the identification of those fetuses that could develop a postnatal neurological impairment.^{3,4,9-12}

The interest in fetal behavior started a long time ago with two-dimensional (2D) ultrasound.¹³⁻¹⁵ Although 2D ultrasonography documents specific fetal movements, only the number of movements was seen, not the quality. More importantly, simultaneous imaging of complex facial movements was impossible using only a 2D ultrasound.^{16,17}

A technique was needed to enable three-dimensional (3D) imaging of fetal facial movements in real-time mode. This technique was introduced a 4D ultrasound (3D in real time).

The first studies on fetal behavior by 4D ultrasound (4D US) validated that this method could help in better understanding of neurosomatic development of the fetus, beginning a new era in the understanding of neurological diseases that start during intrauterine life.^{3,18-21}

Thus, it was suggested that the assessment of fetal behavior in different periods of pregnancy could distinguish between normal and abnormal brain development and early diagnosis of various structural or functional abnormalities.³

For evaluation of fetal behavior, Kurjak et al introduced and described original scoring system called Kurjak's antenatal neurological test (KANET).²²

The parameters used in the test arose from a multicenter survey results conducted earlier on fetal neurobehavioral assessed by 2D ultrasound (2D US) and combined with neonatal signs suggested by Amiel-Tison.^{21,23,24}

Department of Fetal Medicine, Ecomoinhos—School and Clinic of Ultrasonography, Porto Alegre, Brazil

Corresponding Author: Raul Moreira Neto, Ecomoinhos, rua Tobias da Silva 120/501, CEP 90570-020, Porto Alegre, RS, Brazil
Phone: (55)51-3346-2111, e-mail: raul@ecomoinhos.com.br

The primary goal of this pilot study test was to see the feasibility of using KANET in high-risk and low-risk pregnancies in Brazil, to develop and test adequacy of research instruments, assessing the possibility of a full-scale study in a reference hospital and designing a research protocol.

PATIENTS AND METHODS

During a period of almost 2 months, between November and December of 2014, KANET was applied in 51 singleton pregnancies between 28th and 38th weeks of gestation. Gestational age was estimated by the first day of last menstrual period and confirmed by the first ultrasound assessment.

Fetal behavior was assessed by 4D US. All examinations were performed by a single experienced operator using Voluson E8 (GE Healthcare ultrasound, Milwaukee, WI, USA), with volumetric transabdominal 5 MHz transducer.

Each participant gave her informed consent to the study.

The study participants were those who came for an ultrasound exam in a private fetal medicine clinic, in Porto Alegre, Brazil. Were included 17 high-risk and 34 low-risk pregnancies as controls. The inclusion criteria for high-risk pregnancies are presented in the Table 1.

For assessment of fetal neurobehavior, we used a KANET scoring system²² presented in Table 2.

Table 3 shows the interpretation of the KANET scores.

After a systematic 2D measurement of fetal growth and examination of placenta and amniotic fluid volume, the mothers were offered an assessment of fetal behavior by KANET scoring system, using 4D US. The exam was performed while fetuses were awake. Mothers should avoid food, coffee and tea for 2 hours before the examination. The examination maximum time was 30 minutes. According to the score result, fetal behavior was classified into three groups: normal, with total KA-

NET score of 10 to 16, borderline with total score of 6 to 9 and abnormal with total score of £ 5.

The description and the meaning of each specific movement and sign of KANET is described elsewhere.²⁸

The primary outcome was the usefulness of KANET to identify fetuses from high-risk pregnancies at neurological risk. Then we want to compare the scores between the high-risk and low-risk group and, also, compare the KANET results between the high-risk group and subgroups of fetuses. Due to relatively small sample size statistical analysis is at present impossible.

RESULTS

In this prospective longitudinal study, the KANET was applied in 51 singleton pregnancies between 28 and 38 weeks of gestation. There were 17 pregnant women in the high-risk group and 34 pregnant women in low-risk group.

The final results of KANET study, only high-risk patients showed abnormal scores (17.6%). 80.6% of high-risk patients had borderline results while 85.3% of low-risk patients were normal, both statistically significant. Table 4 shows the allocation of fetuses according to KANET.

The comparison of individual parameters for KANET score 0, between fetuses of low and high-risk pregnancies, there was a statistically significant difference for isolated head anteflexion, cranial sutures and head circumference, isolated hand movement or hand to face movements, isolated leg movement and fingers movements. There was no difference between isolated eye blinking , facial movements or mouth opening and gestalt perception of general movements.

On other hand, for KANET score 2, there was significant difference between all the parameters. Table 5 shows the comparison of individual parameters in KANET.

For the three abnormal KANET results (score between 0 and 5), one was related to pregnancy condition (preeclampsia) and two were related to fetal condition (trisomy 13 and Intrauterine growth restriction).

Table 6 presents individual participants assigned to the high-risk group according to prenatal diagnosis.

CONCLUSION

In the last three decades, there have been several attempts to create a diagnostic test for assessment of fetal behavior. Although several were important, none of them was implemented in daily practice.^{22,25-27} KANET is the first prenatal neurological screening test based on 4D US technique and the first results are encouraging.^{13-15,28-30} We believe that KANET can separate serious structural

Table 1: Inclusion criteria

Family history	Previous child with cerebral palsy
Maternal condition	Diabetes mellitus type I and II, thyroid disease, preexistent hypertension, drug abuse, thrombophilia, anemia, epilepsy, fever > 39°C
Pregnancy-related disorders	Gestational diabetes, Rh immunization, threatened preterm labor, preeclampsia, intrauterine infections, viral illness, cholestasis
Fetal condition	Structural and chromosomal abnormalities, polyhydramnion, intrauterine growth restriction, abnormal cardiotocography, biophysical profile and Doppler findings



Table 2: Scoring system of Kurjak antenatal neurodevelopmental test (KANET)

Sign	Score		
	0	1	2
Isolated head anteflexion 	Abrupt	Small range (0-3 times of movements)	Variable in full range, many alteration (>3 times of movements)
Cranial sutures and head circumference 	Overlapping of cranial sutures	Normal cranial sutures with measurement of HC below or above the normal limit (-2 SD) according to GA	Normal cranial sutures with normal measurement of HC according to GA
Isolated eye blinking 	Not present	Not fluent (1-5 times of blinking)	Fluency (>5 times of blinking)
Facial alteration (grimace or tongue expulsion)  or mouth opening (yawning or mouthing) 	Not present	Not fluent (1-5 times of alteration)	Fluency (>5 times of alteration)
Isolated leg movement 	Cramped	Poor repertoire or small in range (0-5 times of movement)	Variable in full range, many alternation (>5 times of movement)
Isolated hand movement  or hand to face movements 	Cramped or abrupt	Poor repertoire or small in range (0-5 times of movement)	Variable in full range, many alternation (>5 times of movements)
Fingers movements 	Unilateral or bilateral clenched fist, (neurological thumb)	Cramped invariable finger movements	Smooth and complex, variable finger movements
Gestalt perception of GMS	Definitely abnormal	Borderline	Normal

Table 3: Interpretation of total score

Total score	Interpretation
0-5	Abnormal
6-9	Borderline
10-16	Normal

Table 4: Interpretation of KANET score in low and high-risk pregnancies

Total score interpretation	High-risk (n = 17)	Low-risk (n = 34)
0-5 Abnormal	3 (17.6%)	0
6-9 Borderline	12 (70.6%)	5 (14.7%)
10-16 Normal	2 (11.8%)	29 (85.3%)

Table 5: Comparison of individual parameters in Kurjak's antenatal neurological test between high-risk and low-risk pregnancies

KANET parameters	Kanet scores		
	0	1	2
<i>Isolated head anteflexion</i>			
High-risk (n:17)	5 (29.4%)	10 (58.9%)	2 (11.7%)
Low-risk (n = 34)	0	16 (47.1%)	18 (52.9%)
<i>Cranial sutures and head circumference</i>			
High-risk (n:17)	3 (17.6%)	6 (35.2%)	8 (47.2%)
Low-risk (n = 34)	0	4 (11.8%)	30 (88.2%)
<i>Isolated eye blinking</i>			
High-risk (n:17)	4 (23.5%)	7 (41.2%)	6 (35.3%)
Low-risk (n = 34)	0	9 (37.5%)	25 (72.5%)
<i>Facial movements (grimace or tongue expulsion) or mouth opening (yawning or mouthing)</i>			
High-risk (n:17)	2 (11.7%)	8 (47.2%)	7 (41.1%)
Low-risk (n = 34)	0	10 (29.4%)	24 (70.6%)
<i>Isolated hand movement or hand to face movements</i>			
High-risk (n:17)	3 (17.6%)	9 (52.9%)	5 (29.5%)
Low-risk (n = 34)	0	9 (37.5)	25 (72.5)
<i>Isolated leg movement</i>			
High-risk (n:17)	4 (23.5%)	10 (58.9%)	3 (17.6%)
Low-risk (n = 34)	0	12 (35.2%)	22 (64.8%)
<i>Fingers movements</i>			
High-risk (n:17)	6 (35.3%)	6 (35.3%)	5 (29.4%)
Low-risk (n = 34)	0	10 (29.4%)	24 (70.6%)
<i>Gestalt perception of general movements</i>			
High-risk (n:17)	2 (11.8%)	9 (52.9%)	6 (35.3%)
Low-risk (n = 34)	0	9 (37.5%)	25 (72.5%)

problems associated with brain injuries. Therefore, our aim is to assess whether KANET really has the potential for prenatal detection of serious neurological problems as well as other causes. Although promising, the concept

Table 6: High-risk study group according to prenatal diagnosis and KANET results

High-risk group	Total	KANET results		
		Normal	Borderline	Abnormal
Preeclampsia	3	0	2	1
Threatened preterm labor	2	0	2	0
Preexistent hypertension	3	1	2	0
Gestational diabetes	2	0	2	0
Anemia	1	1	0	0
Polyhydramnion	1	0	1	0
Intrauterine growth restriction	4	0	3	1
Trisomy 13	1	0	0	1
Total	17	2	12	3

of KANET and its diagnostic value in the evaluation of fetal central nervous system integrity must be confirmed by studies that are underway in several research centers around the world, and we want to continue this research in Brazil as well.

In this pilot study, we saw that there are important differences between the behavior of normal and high-risk fetus, which is in agreement with other studies.^{22,25-27,31-33}

We also found that the vast majority of high-risk pregnancies had borderline KANET. On the other hand, low-risk pregnancies most often presented normal results. But we need to do a bigger research to see the efficiency and effectiveness of KANET. This project was essential to analyze the feasibility of a study on a larger scale and we saw it was realistic and achievable to adapt the protocols, methods and instruments. In addition, logistical problems have been identified and we saw what resources (financial, staff) are needed for a planned study.

We have reached our goal in the study and found that it is possible to do KANET research on a larger scale in Brazil. We also saw the potential to detect and discriminate normal fetal behavior of borderline and abnormal in high-risk pregnancies using KANET.

REFERENCES

1. Rees S, Harding R. Brain development during fetal life: influences of the intra-uterine environment. *Neurosci Lett* 2004;361:111-114.
2. Harrison PJ. The neuropathology of schizophrenia. A critical review of the data and their interpretation. *Brain* 1999;122: 593-624.
3. Prechtl HFR. Qualitative changes of spontaneous movements in fetus and preterm infant are a marker of neurological dysfunction. *Early Hum Dev* 1990;23:151-158.
4. Hadders-Algra M. General movements: a window for early identification of children at high-risk of developmental disorders. *J Pediatr* 2004;145:S12-S18.
5. Cioni G, Prechtl HFR, Ferrari F, Paolicelli PB, Einspieler C, Roversi MF. Which better predicts later outcome in



- full-term infants: Quality of general movements or neurological examination? *Early Hum Dev* 1997;50:71-85.
6. Einspieler C, Prechtl HFR, Bos AF, Ferrari F, Cioni G. Prechtl's method on the qualitative assessment of general movements in preterm, term and young infants. London: Mac Keith Press 2004.
 7. Einspieler C, Prechtl HFR. Prechtl's assessment of general movements: a diagnostic tool for the functional assessment of the young nervous system. *Ment Retard Dev Disabil Res Rev* 2005;11:61-67.
 8. Strijbis EMM, Oudman I, Van Essen P, MacLennan AH. Cerebral palsy and application of the international criteria for acute intrapartum hypoxia. *Obstet Gynecol* 2006;107:1357-1365.
 9. Amiel-Tison A, Gosselin J, Kurjak A. Neurosonography in the second half of fetal life: a neonatologist's point of view. *J Perinat Med* 2006;34:437-446.
 10. Groen SE, de Blecourt AC, Postema K, Hadders-Algra M. General movements in early infancy predict neuromotor development at 9-12 years of age. *Dev Med Child Neurol* 2005;47:731-738.
 11. Hepper PG. Fetal habituation: another Pandora's box? *Dev Med Child Neurol* 1997;39:274-278.
 12. Stanojevic M, Kurjak A, Andonotopo W. Assessment of fetal to neonatal behavioral continuity by 4D ultrasonography. *Ultrasound Obstet Gynecol* 2006;28:360.
 13. Nijhuis JG, Prechtl HF, Martin CB Jr, Bots RS. Are there behavioural states in the human fetus? *Early Hum Dev* 1982;6:177-195.
 14. Horimoto N, Koyanagi T, Maeda H, Satoh S, Takashima T, Minami T, Nakano H. Can brain impairment be detected by in utero behavioural patterns? *Arch Dis Child* 1993;69 (1 Spec No):3-8.
 15. Prechtl HFR, Einspieler C. Is neurological assessment of the fetus possible? *Eur J Obstet Gynecol Reprod Biol* 1997;75: 81-84.
 16. Arabin B. Two-dimensional real-time ultrasound in the assessment of fetal activity in single and multiple pregnancies. *Ultrasound Rev Obstet Gynecol* 2004;4:37.
 17. Sepulveda W, Mangiamarchi M. Fetal yawning. *Ultrasound Obstet Gynecol* 1995;5:57.
 18. Stanojevic M, Kurjak A, Salihagic-Kadic A, Vasilj O, Miskovic B, Shaddad AN, Ahmed B, et al. Neurobehavioral continuity from fetus to neonate. *J Perinat Med* 2011;39:171-177.
 19. Kuno A, Akiyama M, Yamashiro C, Tanaka H, Yanagihara T, Hata T. Three-dimensional sonographic assessment of fetal behavior in the early second trimester of pregnancy. *J Ultrasound Med* 2011;20:1271-1275.
 20. Horimoto N, Koyanagi T, Satoh S, Yoshizato T, Nakano H. Fetal eye movement assessed with real-time ultrasonography: are there rapid and slow eye movements? *Am J Obstet Gynecol* 1989;163:1480-1484.
 21. Yigiter AB, Kavak ZN. Normal standards of fetal behavior assessed by four-dimensional sonography. *J Matern Fetal Neonatal Med* 2006;19:707-721.
 22. Kurjak A, Miskovic B, Stanojevic M, Amiel-Tison C, Ahmed B, Azumendi G, Vasilj O, Andonotopo W, Turudic T, Salihagic-Kadic A. New scoring system for fetal neurobehavior assessed by three and four-dimensional sonography. *J Perinat Med* 2008;36:73-81.
 23. Amiel-Tison C. Update of the Amiel-Tison neurological assessment for the term neonate or at 40 weeks corrected age. *Pediatr Neurol* 2002;27:196-212.
 24. Kurjak A, Andonotopo W, Hafner T, Salihagic A, Standojevic M, Azumendi G, et al. Normal standards for fetal neurobehavioral developments—longitudinal quantification by four-dimensional sonography. *J Perinat Med* 2006;34:56-65.
 25. Kurjak A, Abo-Yaqoub S, Stanojevic M, Yigiter AB, Vasilj O, Lebit D, et al. The potential of 4D sonography in the assessment of fetal neurobehavior: Multicentric study in high-risk pregnancies. *J Perinat Med* 2010;38(1):77-82.
 26. Miskovic B, Vasilj O, Stanojevic M, Ivankovic D, Kerner M, Tikvica A. The comparison of fetal behavior in high-risk and normal pregnancies assessed by four-dimensional ultrasound. *J Matern Fetal Neonatal Med* 2010 Dec;23(12): 1461-1467.
 27. Talic A, Kurjak A, Ahmed B, Stanojevic M, Predojevic M, Salihagic Kadic A, Di Renzo GC. The potential of 4D sonography in the assessment of fetal behavior in high-risk pregnancies. *J Matern Fetal Neonatal Med* 2011 Jul;24(7): 948-954.
 28. Kurjak A, Ahmed B, Abo-Yaguab S, Younis M, Saleh H, Shaddad AN, et al. An attempt to introduce neurological test for fetus based on 3D and 4D sonography. *Donald School Journal of Ultrasound in Obstetrics and Gynecology* 2008; 2:29-44.
 29. Kurjak A, Tikvica Luetic A, Stanojevic M, Talic A, Zalud I, Al-Noobi M, et al. Further experience in the clinical assessment of fetal neurobehavior. *Donald School J Ultrasound in Obstet and Gynecol* 2010;4:59-71.
 30. Morokuma S, Fukushima K, Yumoto Y, Uchimura M, Fujiwara A, Matsumoto M, Satoh S, Nakano H. Simplified ultrasound screening for fetal brain function based on behavioral pattern. *Early Hum Dev* 2007;83:177-1781.
 31. Predojević M, Stanojević M, Vasilj O, Kadić AS. Postnatal neurological evaluation of a fetus and newborn from pregnancy complicated with IUGR and fetal hypoxemia. *J Matern Fetal Neonatal Med* 2011 May;24(5):764-767.
 32. Athanasiadis AP, Mikos T, Tambakoudis GP, Theodoridis TD, Papastergiou M, et al. Neurodevelopmental fetal assessment using KANET scoring system in low and high risk pregnancies. *J Matern Fetal Neonatal Med* 2013;26:363-368.
 33. Abo-Yaqoub S, Kurjak A, Mohammed AB, Shadad A, Abdel-Maaboud M. The role of 4-D ultrasonography in prenatal assessment of fetal neurobehaviour and prediction of neurological outcome. *J Matern Fetal Neonatal Med* 2012;25: 231-236.