157 EFFECTS OF ACUTE HYPOXIA ON THE CELLULAR ELECTRICAL ACTIVITY OF NEONATAL AND ADULT PURKINJE FIBERS (PF). Steven M. Yabek, Rinya Kato, Bramah N. Singh, Dept. of Pediatrics, University of New Mexico, Albuquerque, NM.

We previously described that action potential (AP) parameters from neonatal (N) ventricular myocardium are more resistant than adult (A) APs to acute hypoxia (H). In this study, we used standard microelectrode techniques to evaluate the effects of acute H (pO₂=30 torr) on A (n=10) and N (3-12 days old)(n=11) canine PF APs and PF arrhythmogenesis. PFs were paced at 1 Hz. Compared to A, control AP from N PF had significantly (s)(p<0.05) lower values for AP amplitude (APA), Vmax, AP duration at 50% repolarization (APD50), APD90 and effective refractory period (ERP). After 30 min. of acute hypoxic superfusion, N PF showed s greater reductions than A PF in Vmax (-20% vs -9%), APD50 (-18% vs -8%) and APD90 (-13% vs -4%). Changes in APA, maximum diastolic potential (MDP) and ERP did not differ. All N AP parameters (except MDP) were s reduced after H. Reoxygenation for 30 min. partially restored N AP indices. When pacing was stopped during control, no A or N PF had spontaneous activity (sa) for 15 sec. H produced sa in 0 of 10 A and 2 of 11 N PF. The sa increased during early reoxygenation (1 of 10 in A; 4 of 11 in N) but was abolished in all after 30 min. The H and reoxygenation related ectopic sa was due to both enhanced automaticity and reentry, the latter probably induced by the decreases in Vmax (conduction velocity), APD and ERP. In summary, N PF have an enhanced sensitivity to acute H. The arrhythmogenic effects of H in-vitro may explain the H and reperfusion related arrhythmias seen clinically in some children.

158 AMIODARONE (AM) ADVERSELY AFFECTS SINUS NODE (SN) FUNCTION IN VITRO. Steven M. Yabek, Rinya Kato, Bramah N. Singh, Department of Pediatrics, University of New Mexico, Albuquerque & Wadsworth VA Hospital, L.A.

AM is being used more frequently to control "resistant" childhood arrhythmias. As a Class III agent, AM supposedly acts by uniformly prolonging action potential (AP) duration and refractoriness. Because sinus slowing is observed clinically, we evaluated the acute effects of 0.68-34µg/ml AM on SN pacemaker activity using rabbit SN preparations and standard microelectrode techniques. Data were collected only from true SN pacemaker cells impaled continuously throughout the study. AM produced significant (p<0.05) concentration dependent decreases in spontaneous sinus rate (-6% at 6.8µg/ml; -12% at 34µg/ml), AP amplitude (-15% at 34µg/ml) and rate of phase 4 diastolic depolarization (-14%, -18%, -27% at 0.68, 6.8 & 34µg/ml respectively). Maximum diastolic & threshold potentials and Vmax also decreased significantly. Since AP duration did not increase, the fall in spontaneous rate was related solely to AM induced decreases in automaticity. Corrected SNRT and SACT were increased markedly by AM. In 4 preparations, SN exit or entrance block developed on prolonged exposure. The AM induced changes could not be reversed despite superfusion in AM-free medium for up to 1.5 hrs. Rather, SN function deteriorated further. AMs Class III actions cannot explain its effects on the SN. Since the automaticity and upstroke of SN pacemaker activity are dependent upon the slow inward Ca⁺⁺ current (i_{si}), AM may also possess Class IV (verapamil-like) actions or marked antiadrenergic activity in-vitro.

159 INDOMETHACIN (I) THERAPY IN PREMATURE INFANTS OF OLD POSTNATAL AGE (>8 WEEKS). T.F. Yeh, B. Achanti, J. Stubb, S. Pyati, R.S. Pildes, Cook County Hosp., Dept. of Ped., Univ. of Ill., Chicago, Ill.

Previous studies have shown a positive correlation between plasma I level and ductus closure and between clearance of plasma I and postn. age; it is suggested that failure of ductus response to I in some infants of old postnatal age may have been due to low plasma I. To test this hypothesis, 5 infants with postn. age >8 wks were given iv I 0.3 mg/kg q 6 hr for 4 doses in an attempt to maintain high plasma I. All infants had sign. PDA (clinical cardiovascular distress score (CVD) >3 and Echo LA/AO >1.3) at time of study. The mean±S.D. birth wt was 969±209 gm; Gest. age 29±1.4 wks, Postn. age 65.2±4.3 days. Plasma samples for I assay were obtained at ½, 4, 8, 16, 24 hrs after 1st dose.

	½	4	8	16	24
Plasma I (ng/ml)	828±81	350±90	590±102	570±88	370±96
Echo:LA/AO	1.48±0.29	1.51±0.30	1.50±0.27	1.49±0.32	1.49±0.28
CVD score	5.2±1.2	4.8±1.1	5.1±1.2	5.1±1.0	5.0±1.1

None of the infants responded to I with ductus closure in spite of maintenance of plasma I concentration above presumptive therapeutic level (250 ng/ml at 24 hr post-dosing). There was no significant changes in LA/AO and CVD score. However, significant decreases in urine output (from 3.1±0.7 to 2.1±0.6 ml/kg/h, p <0.01) and FENa (1.5±0.4 to 1.0±0.3%, p <0.05) were seen after the study. This study suggests that failure to I therapy in infant of old postn. age may not relate to low plasma I level and that a primary histological anomaly of the ductus independent of PG might be present in these infants.

160 RAPID ATRIAL PACING-INDUCED INFRA-HIS CONDUCTION BLOCK Ming-Lon Young, Henry Gelband, Augustin Castellanos, Grace S. Wolff, Univ of Miami, Jackson Memorial Hosp, Miami.

Nine out of 285 (3%) pediatric patients (pts) were found to have pacing-induced infra-His conduction block (B). In 2 pts it occurred following a preceding long cycle as well as a preceding short-to-long sequence cycle length (CL) change. In one of them the infra-His B initiated a short sequence of 2:1 infra-His B. In 5 pts it occurred with an abrupt shortening of the coupling H-H (in one pt it simulated type II 2nd-degree AV B). In 2 pts the pacing-induced infra-His B simulated type II 2nd-degree AV block but the initiating events were not recorded. Rapid atrial pacing-induced infra-His B in children is a rare phenomenon. In many cases it is associated with pacing-induced infra-His conduction delay. The occurrence of either is determined by: (1) a preceding long cycle as well as a preceding short-to-long sequence CL change that can lengthen the refractoriness of the His-Purkinje system (HPS), (2) an abrupt shortening of the coupling H-H interval that exceeds the refractory period of the HPS, (3) a first impulse that results in an infra-His B initiating a series of 2:1 infra-His B due to a self-perpetuation sequence of events, (4) a 1st impulse that results in infra-His conduction delay initiating a series of 1:1 infra-His delay due to repetitive retrograde concealed conduction.

161 REVERSED ALTERNATING WENCKEBACH PERIODICITY Ming-Lon Young, Henry Gelband, Augustin Castellanos, Grace S. Wolff, Univ of Miami, Jackson Memorial Hosp, Dept of Pedi Cardiology, Miami.

Alternating Wenckebach periodicity (AWP) is defined as an pattern of 2:1 AV block associated with progressive lengthening of the conducted impulses with termination in a higher degree of block. We found another phenomenon in 5 children during atrial pacing which is the reverse of this pattern (RAWP) and has not been previously reported: that is, during an underlying pattern of 2:1 AV block, there is progressive shortening of the conducted impulses with termination in a lower degree of block. The mechanism can be explained by a one-level block model. It is formed by a series of self-perpetuating events: (1) The 1st beat of the RAWP is coupled to a preceding impulse with a short coupling interval (CI) and results in a long (the longest) delay. (2) With a long preceding delay the 2nd beat of the RAWP has a short (the shortest) CI falling within the effective refractory period of the AVN (ERP-AVN) and results in block. (3) Therefore the 3rd beat is actually coupled to the 1st beat with a long CI and results in a shorter delay. (4) The 4th beat also has long CI, which, if still shorter than the ERP-AVN results in a sequence of 2:1 block in which the conducted impulses showing progressive shortening. This continues until CI of the even numbered beat exceeds the ERP-AVN and, thereby, conducts.

162 ALTERNATING WENCKEBACH PERIODICITY AND MULTILEVEL AV BLOCK Ming-Lon Young, Henry Gelband, Grace S. Wolff, University of Miami, Jackson Memorial Hospital, Department of Pediatric Cardiology, Miami, Florida.

Six pediatric patients (pts) exhibiting atrial pacing-induced repetitive block (B) are presented. Analysis of the conduction patterns satisfy the requisites for multilevel AV B. 1) DOCUMENTED 2 LEVEL B: Two pts exhibited an upper level AVN Wenckebach periodicity (WP) and a lower level 2:1 infra-His B. 2) DEDUCED 2 LEVEL B IN THE AVN: Two pts had 2:1 AVN B terminated by 3:1 B (alternating WP). With group 1 serving as a model, it is postulated that in this group the 2 sites of B are within the AVN: an upper level AVN WP and a lower level 2:1 AVN B. One pt had an AVN WP terminated by 3:1 B. This was followed by 2 series of 2:1 B, suggesting simultaneous WP at 2 AVN levels. 3) DEDUCED 3 LEVEL B IN THE AVN: One pt had 2:1 AVN B with progressive prolongation of the conducted AH terminating in 2 series of 3:1 B in which the AH following the 1st 2 blocked atrial beats was not the shortest while that following the 2nd was. With previous groups serving as models, it is postulated that there is a 3 level B inside the AVN: an upper level AVN WP, a middle level AVN WP of higher degree B and a lower level 2:1 AVN B. In summary, we have for the first time, described multilevel B within the AVN in the pediatric pt.