During subsequent post-ischemic hypoperfusion, in both 
C0-reactivity and autoregulation are abolished.

2. During reactive hyperemia, cerebral vessels are paralyzed, and 
increased metabolic demand of the tissue and therefore 
oxygen availability is in misrelationship to an in­
crease of cerebral venous blood increases (luxury perfusion).

ated with disturbances of flow regulation. During reac­
may result in relative cerebral hypoxia.

2. During post-ischemic hypoperfusion, the decreased 
requirements of the tissue, and oxygen content 
in reactive hyperemia, oxygen availability exceeds the 
producing metabolism and blood flow are uncoupled:
followed by a phase of reduced blood circulation (post­
phases, energy-
producing metabolism and blood flow are uncoupled: 
in reactive hyperemia, oxygen availability exceeds the 
requirements of the tissue, and oxygen content 
of cerebral venous blood increases (luxury perfusion).

12. Post-ischemic hypoperfusion, the decreased oxygen availability is in misrelationship to an 
increased metabolic demand of the tissue and therefore 
may result in relative cerebral hypoxia.

2. The hemodynamic changes are caused by or associ­
ated with disturbances of flow regulation. During reactive 
hyperemia, cerebral vessels are paralyzed, and both 
C02-reactivity and autoregulation are abolished. 
During subsequent post-ischemic hypoperfusion, in

contrast, vascular tone is increased although C02-reactivity is abolished, and a rise of blood pressure now 
causes an autoregulatory constriction of the resistance 
vessels. In consequence, blood flow in this period of 
hypoperfusion cannot be improved by either increasing 
blood pressure or increasing arterial or tissue pCO2.

3. The hemodynamic changes observed during post­
 ischemic hypoperfusion resemble the pharmacological 
effect of indomethacin, an inhibitor of prostaglandin 
synthesis, which also reduces blood flow and abolishes 
C02-reactivity without interfering with autoregula­
tion. Prostaglandins have been shown to play an im­
portant role in the hemodynamic balance of local blood 
circulation, whereby prostacyclin (PGI2) and thromboxane A2 (TXA2) are mainly involved. The 
respective precursors are the cyclic endoperoxides PGG2 and PGH2, that are synthetized from arachidonic 
acids by cyclo-oxygenases. These endoperoxides 
are transformed in the vascular wall by prostacyclin synthetase into PGI2 and thromboxane synthetase into TXA2.

4. Prostaglandines are transformed in the vascular wall by prostacyclin synthetase into PGI2 and thromboxane synthetase into TXA2. This 
compound reverses the reduction of cerebral blood flow and restores

SUMMARY In normothermic cats under light barbiturate anesthesia, cerebral blood flow was arrested 
for one hour by intrathoracic occlusion of the innominate, the left subclavian, and both mammarian 
arteries. Recirculation of the brain after ischemia resulted in reactive hyperemia, followed by a decrease of 
blood flow to about 70% of control (post-ischemic hypoperfusion). During postischemic hypoperfusion, 
C02-reactivity was completely abolished. Intravenous infusion of prostacyclin 2 hours after ischemia (1.8 
µg/kg/min) decreased systemic arterial blood pressure and reduced platelet aggregability but did not 
improve cerebral blood flow, did not restore C02-reactivity, and did not influence postischemic changes of 
blood coagulation. It is concluded that prostacyclin deficiency is not or not the only reason for the 
development of post-ischemic hypoperfusion and the associated disturbance of flow regulation.

Address correspondence to: W. van den Kerkoff, Max-Planck-Institut für neurologische Forschung, Abtei­lung für experimentelle Neurologie, und Lehrstuhl Innere Medizin II, Universität Köln, Ostmerheimer Str. 200 D-5000 Köln 91 (Merheim), Republic of West Germany.

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From the Max-Planck-Institut für neurologische Forschung, Abtei­lung für experimentelle Neurologie, und Lehrstuhl Innere Medizin II, Universität Köln, Ostmerheimer Str. 200 D-5000 Köln 91 (Merheim), Republic of West Germany.
CO₂-reactivity. In view of these findings, the question arose whether post-ischemic hypoperfusion and the associated disturbance of CO₂-reactivity can be reversed by therapeutic application of prostacyclin. Post-ischemic hypoperfusion is frequently associated with regional disturbances of metabolic activity and of the blood coagulation system. These effects, therefore, were also studied. In this communication the hemodynamic and hemostatic, and in an accompanying paper the metabolic observations will be described.

Material and Methods

Thirteen adult cats of both sexes, weighing 2–3.5 kg, were used. The animals were anesthetized with pentobarbital (Nembutal®, 30 mg/kg intraperitoneally). Following tracheotomy, the animals were immobilized by pancuronium bromide (Pancuronium®, Organon, FRG) and mechanically ventilated with a Harvard respirator. Arterial pCO₂ was maintained close to 30 mm Hg and arterial pO₂ above 100 mm Hg by appropriate adjustment of tidal volume and oxygen content of the inhaled air. Tidal CO₂ was continuously monitored using a CO₂-analyzer (URAS, Hartmann and Braun, Frankfurt, FRG). Body temperature was maintained at 37°C with a thermocontrolled heating pad.

Catheters were placed into the following vessels: into the innominate artery via the left brachial artery for estimation of cerebral blood flow by bolus injection of 133Xenon; into the left femoral artery for monitoring the systemic arterial blood pressure and for withdrawal of arterial blood samples; into both femoral veins for drug infusion and stabilization of systemic blood pressure by withdrawal or reinfusion of blood, if necessary.

Blood flow of the total brain was completely arrested for one hour by intratracheal occlusion of the innominate, the left subclavian and both mammalian arteries. Collateral blood supply via the ascending spinal arteries was prevented by lowering the systolic blood pressure below 80 mm Hg with trimethaphan-camphorsulfonate (Arfonad®) and — if necessary — by withdrawal of blood. The completeness of ischemia was controlled by recording the clearance of 133Xenon, that was injected immediately before the occlusion of the arteries. Ischemia was considered to be complete when radioactivity over the head decreased by less than 10% during the total length of ischemia.

In order to promote blood recirculation after ischemia, the animals were submitted to the following procedure. Ten minutes before the end of the ischemia, 30 ml Tris buffer (Sterofundin-Tris®, 15 mVal/l) were infused at a rate of 1 ml/min for equilibration of the acid-base balance during the early recirculation period. Five minutes before recirculation, infusion of 10 ml/kg 20% mannitol (Osmosteril®), was started at a rate of 1 ml/min to reduce brain swelling which regularly occurs during reperfusion. Immediately before recirculation, systolic blood pressure was abruptly elevated to more than 180 mm Hg by intravenous infusion of norfenedrine (Novadril®), and the vascular clamps were removed from the innominate and subclavian arteries at the peak of the pressure pulse. Blood reperfusion of the ischemic brain resulted in a steep increase of arterial pCO₂, as evidenced by the rise of tidal CO₂. Respiration rate and volume, therefore, were increased until tidal pCO₂ returned to normal values. Arterial blood samples were taken in short intervals (5–10 min) to control buffer infusion and ventilation rate until acid-base balance and arterial blood gases returned to normal.

As a specific therapy, prostacyclin (1.8 μg/kg/min) was applied in 6 cats by continuous intravenous infusion over 60 min after the second hour of recirculation (PGI₂-treated group). The drug was freshly prepared from a stock solution (1 mg prostacyclin in 1 ml glycine buffer, pH 10.5, 4°C) by dilution 1:5 with cold 15 mVal Tris buffer immediately before the beginning of infusion. Infusion rate of PGI₂ was temporarily reduced when the systemic blood pressure fell below 130 mm Hg. Seven other cats received an infusion of the solvent alone (placebo group).

Before ischemia and after 3 hours of reperfusion, i.e. at the end of the 1 hour's infusion of PGI₂ or the solvent, respectively, a battery of blood coagulation and platelet function tests were carried out. Global coagulation tests such as PTT, TT, and Quick, the factors V, X, XII and the concentration of fibrinogen were measured by use of commercially available test reagents (Boehringer, Mannheim; Behring-Werke, Marburg, FRG), whereby the clotting time was measured with a coagulometer of Schnittger and Gross (Amelung, Lemgo Brake, FRG).

Plasminogen and AT III-activity were measured with a chromogenic substrate assay (Kabi, Stockholm, Sweden), blood platelets with a cell counter (Type 123, Analys Instruments, Stockholm, Sweden), and platelet aggregation ratio (PAR) was evaluated according to the method of Wu and Hoak. ADP-induced platelet aggregation was measured in heparinized whole blood samples immediately after withdrawal with an electronic aggregometer.

Cerebral blood flow was measured before and at various intervals after ischemia using the intraarterial 133Xenon injection technique. For each flow estimation, one mCi 133Xenon (dissolved in about 0.2 ml Ringer solution) was injected as a bolus into the innominate artery, and the 133Xenon clearance from the brain was monitored by an extracranial collimated scintillation detector placed over the exposed skull. Muscles and tongue were shielded with lead to avoid extracerebral contamination of the clearance curves. Clearance curves were evaluated by the 2 minutes' slope index.

CO₂-reactivity of cerebral blood flow was estimated before ischemia and 3 hours after the beginning of recirculation by ventilating the animal with a gas mixture containing 6% CO₂, 21% oxygen, and the rest nitrogen.

The electrocorticogram was recorded from the sensory motor area of the right hemisphere with bipolar
silverball electrodes. The same region was also stimulated with single rectangular pulses (0.3 ms, 10 V), and the evoked pyramidal response was recorded at the bulbar level using a stereotactically implanted concentric needle electrode.

Restitution of electrophysiological function after ischemia was classified 3 hours after the beginning of recirculation as follows:

**Recovery** was defined as normalization of the pyramidal response, accompanied by the beginning recovery of EEG. **Partial recovery** was defined as a partial return of the pyramidal response without recovery of EEG. **No recovery** indicated absence of any neurophysiological functions.

At the end of the experiments the brains were frozen in situ with liquid nitrogen. The brains were sawed in slices of 0.5 cm thickness, and processed for regional biochemical evaluation of NADH, glucose and ATP, as described elsewhere.¹

### Results

Two groups of animals were compared. In one group (7 cats, placebo group) global complete cerebral ischemia of one hour was followed by three hours' blood recirculation using a standardized therapy for prevention of no-reflow (see Methods). In another group of animals (6 cats, PGI₂-treated group) the same therapy was supplemented by continuous intravenous infusion of prostacyclin (1.8 μg/kg/min), starting two hours after beginning of recirculation and continuing until the end of the experiments after 3 hours.

#### General Physiological Parameters

The data obtained are summarized in table 1: In both groups, the acid-base status of the blood, blood gases and hematocrit remained stable throughout the experiments. During recirculation, the mean arterial blood pressure, after a period of induced hypertension, decreased to about 85 mm Hg after one hour, and subsequently stabilized between 100 and 150 mm Hg after 2 hours of recirculation. In the prostacyclin-treated group, mean arterial blood pressure fell to about 80 mm Hg during drug administration, indicating a general peripheral vasodilating effect.

### Cerebral Blood Flow

In both groups, mean cerebral blood flow was about 40 ml/100 g/min before ischemia (table 1). After vascular occlusion blood flow completely stopped, as confirmed by the absence of ¹³³Xenon-clearance. Immediately after ischemia, reactive hyperemia developed, the degree of which depended on the individual blood pressure. One hour after the onset of recirculation, blood flow stabilized between 30 and 35 ml/100 g/min, i.e. about 25% below the control level. During prostacyclin infusion, blood flow further decreased to 27 ± 3 ml/100 g/min, but this value was not significantly different from that of the untreated animals in which blood flow after 3 hours' recirculation was 29 ± 6 ml/100 g/min. This finding does not exclude that prostacyclin slightly reduced cerebrovascular resistance because blood pressure was lower than in the untreated animals.

**CO₂-reactivity of blood flow** was tested before and three hours after cerebral ischemia (fig. 1). Both before and after ischemia, the ventilation of the animals with 6% CO₂ caused an increase of arterial pCO₂ from 30 to about 60 mm Hg. Before ischemia, hypercapnia led to an increase of blood flow up to 350%. Three hours after recirculation, CO₂-reactivity was completely abolished in both the placebo and prostacyclin-treated group. Prostacyclin, in consequence, was not

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**Table 1** General Physiological Parameters

<table>
<thead>
<tr>
<th></th>
<th>Control</th>
<th>1 hour</th>
<th>2 hours</th>
<th>3 hours</th>
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<tbody>
<tr>
<td><strong>Placebo group</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>pH</td>
<td>7.38 ± 0.02</td>
<td>7.45 ± 0.32</td>
<td>7.43 ± 0.27</td>
<td>7.43 ± 0.32</td>
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<tr>
<td>pCO₂ (mm Hg)</td>
<td>31.0 ± 1.0</td>
<td>30.6 ± 2.4</td>
<td>31.7 ± 2.5</td>
<td>32.3 ± 2.7</td>
</tr>
<tr>
<td>pO₂ (mm Hg)</td>
<td>119 ± 10</td>
<td>146 ± 32</td>
<td>142 ± 30</td>
<td>203 ± 37</td>
</tr>
<tr>
<td>HCO₃⁻ (mmol/l)</td>
<td>19.2 ± 0.8</td>
<td>22.5 ± 1.6</td>
<td>21.8 ± 1.3</td>
<td>22.0 ± 1.2</td>
</tr>
<tr>
<td>Hct (vol%)</td>
<td>35.6 ± 2.8</td>
<td>33.4 ± 2.3</td>
<td>33.1 ± 2.5</td>
<td>34.5 ± 2.3</td>
</tr>
<tr>
<td>SAP (mm Hg)</td>
<td>137.0 ± 6.2</td>
<td>84.6 ± 9.5</td>
<td>101.5 ± 6.6</td>
<td>112.9 ± 9.9</td>
</tr>
<tr>
<td>CBF (ml/100 g/min)</td>
<td>38.5 ± 2.1</td>
<td>30.2 ± 4.3</td>
<td>27.5 ± 4.0</td>
<td>28.7 ± 5.8</td>
</tr>
<tr>
<td><strong>PGI₂-treated group</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>pH</td>
<td>7.36 ± 0.01</td>
<td>7.36 ± 0.04</td>
<td>7.40 ± 0.05</td>
<td>7.41 ± 0.03</td>
</tr>
<tr>
<td>pCO₂ (mm Hg)</td>
<td>30.5 ± 0.5</td>
<td>33.3 ± 3.8</td>
<td>31.3 ± 6.8</td>
<td>32.5 ± 1.9</td>
</tr>
<tr>
<td>pO₂ (mm Hg)</td>
<td>121 ± 13</td>
<td>184 ± 28</td>
<td>270 ± 83</td>
<td>209 ± 21</td>
</tr>
<tr>
<td>HCO₃⁻ (mmol/l)</td>
<td>18.5 ± 0.6</td>
<td>19.5 ± 1.4</td>
<td>20.6 ± 2.0</td>
<td>21.2 ± 0.9</td>
</tr>
<tr>
<td>Hct (vol%)</td>
<td>29.8 ± 1.6</td>
<td>30.7 ± 3.8</td>
<td>31.3 ± 8.6</td>
<td>27.5 ± 2.6</td>
</tr>
<tr>
<td>SAP (mm Hg)</td>
<td>120.6 ± 8.3</td>
<td>85.0 ± 7.3</td>
<td>115.6 ± 15.0</td>
<td>79.7 ± 6.9</td>
</tr>
<tr>
<td>CBF (ml/100 g/min)</td>
<td>40.5 ± 2.6</td>
<td>34.4 ± 4.7</td>
<td>31.6 ± 7.0</td>
<td>26.5 ± 3.1</td>
</tr>
</tbody>
</table>

General physiological parameters before and at various recirculation times following 1 hour complete ischemia of the cat brain. PGI₂ was infused between 2 and 3 hours after ischemia.

Values are means ± SEM.
able to reverse the post-ischemic suppression of CO$_2$-reactivity.

Electrophysiological Function

Electrophysiological suppression and recovery after ischemia was evaluated by recording spontaneous ECoG activity, and the pyramidal response following electrical stimulation of the motor cortex (fig. 2). During ischemia, ECoG activity ceased within 15 sec and the pyramidal response after 3–5 min. After restoration of blood flow, in 5 out of 13 animals pyramidal response began to reappear within 10 min, and initial signs of electro-cortical activity were recorded after 1–2 hours (functional recovery). In 5 animals, only the pyramidal response but not the ECoG reappeared after 30–60 min (partial recovery), and in the remaining 3 animals, electrophysiological recovery was absent or a secondary suppression of electrophysiological functions occurred (no recovery). There was no difference in the incidence of no recovery between treated and untreated animals. This is not surprising because prostacyclin infusion was started 2 hours after the beginning of recirculation, i.e., at a time when a clear distinction between animals with and without recovery could already be made. However, there was also no influence of prostacyclin infusion on the progression of recovery in those animals in which electrophysiological signs of function had returned.

The blood coagulation data are shown in table 2. In both groups global coagulation times, and coagulation times for the factors V, X, XII increased after ischemia and fibrinogen concentration decreased significantly and similarly. An increased activation of the fibrinolytic system was also evident in both groups with a significant and similar fall of plasminogen from 0.94 ± 0.7 to 0.57 ± 0.21 IU/ml in the placebo group ($p < 0.01$) and from 0.98 ± 0.14 to 0.43 ± 0.09 IU/ml in the PGI$_2$-treated group ($p < 0.01$).

A significant reduction of active AT III, indicating an increased thrombin turnover with formation of inactive AT III-thrombin-complexes, was also measured in both groups ($p < 0.01$). The only difference between treated and untreated animals concerned platelet aggregability (fig. 3). During PGI$_2$-infusion platelet aggregation ratio (PAR) significantly increased in both

![Figure 1. CO$_2$ reactivity of cerebral blood flow before and 3 hours after complete cerebro-circulatory arrest of 60 min. Above: untreated animals, below: animals treated with prostacyclin (1.8 µg/kg/min intravenously). Arterial pCO$_2$ was increased by ventilating the animals with 6% CO$_2$.](image)

![Figure 2. Electrocorticogram and pyramidal response following electrical stimulation of the motor cortex before (control) and 3 hours after total cerebro-circulatory arrest of 60 min. Recordings are from animals with different degrees of functional disturbances after ischemia.](image)
TABLE 2 Coagulation Parameters

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Placebo group</th>
<th>PGI₂-treated group</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Control</td>
<td>3 hours</td>
</tr>
<tr>
<td>Prothrombin time (sec)</td>
<td>11.5 ± 0.6</td>
<td>19.5 ± 3.3</td>
</tr>
<tr>
<td>PTT (sec)</td>
<td>18.3 ± 2.1</td>
<td>177.8 ± 54.0</td>
</tr>
<tr>
<td>TT (sec)</td>
<td>15.0 ± 1.1</td>
<td>145.4 ± 64.1</td>
</tr>
<tr>
<td>Fibrinogen (mg/dl)</td>
<td>151 ± 32</td>
<td>109 ± 30</td>
</tr>
<tr>
<td>Factor V (sec)</td>
<td>17.5 ± 1.2</td>
<td>20.0 ± 1.5</td>
</tr>
<tr>
<td>Factor X (sec)</td>
<td>39.6 ± 0.8</td>
<td>50.1 ± 3.6</td>
</tr>
<tr>
<td>Factor XII (sec)</td>
<td>28.9 ± 2.9</td>
<td>40.7 ± 2.8</td>
</tr>
<tr>
<td>Plasminogen (CTA-U/ml)</td>
<td>0.94 ± 0.70</td>
<td>0.57 ± 0.21</td>
</tr>
<tr>
<td>AT III (IU/ml)</td>
<td>17.0 ± 2.2</td>
<td>11.5 ± 1.9</td>
</tr>
<tr>
<td>PAR % venous</td>
<td>0.87 ± 0.06</td>
<td>0.75 ± 0.17</td>
</tr>
<tr>
<td>PAR % arterial</td>
<td>0.97 ± 0.03</td>
<td>0.85 ± 0.15</td>
</tr>
</tbody>
</table>

Changes of blood coagulation parameters, fibrinolysis and platelet aggregation ratio (PAR according to Wu and Hoak21) before and 3 hours following 1 hour complete ischemia. PGI₂ was infused between 2 and 3 hours after ischemia. Values are means ± SEM.

the arterial and venous blood, when compared with the preinfusion value (p < 0.05). In the placebo group, on the other hand, platelet aggregation ratio fell. This indicates an increase in in-vivo platelet aggregation during solvent (glycine/Tris buffer 1:5) infusion in contrast to the platelet disaggregating effect of PGI₂.

Discussion

The present experiment was designed in order to test the hypothesis that a reduction of PGI₂ may be responsible for the disturbance of the CO₂-reactivity and the decrease of CBF observed during the phase of post-ischemic hyperperfusion following a period of prolonged cerebro-circulatory arrest. This hypothesis was derived from the hemodynamic similarities between post-ischemic hyperperfusion and the reduction of blood flow by indomethacin which causes an inhibition of prostaglandin synthesis and which is reversed by systemic application of prostacyclin.15

During ischemia, arachidonic acid accumulates in the brain tissue and during the early recirculation phase triggers a shortlasting burst of production of prostaglandins via prostaglandin endoperoxides.15-27 These endoperoxides have a free radical character and may be able to inactivate their respective enzymes.28-30 The preferential inactivation of prostacyclin synthetase can be explained by its location in the wall of cerebral vessels. Thromboxane A₂ synthetase, on the other hand, is less affected because only a small fraction of total platelet content resides in the brain during ischemia. This could lead to an imbalance of vasoactive prostaglandins: the reduced synthesis of the vasodilating and platelet disaggregating PGI₂ in the vessel wall may cause a relative increase of the vasoconstricting and platelet aggregating TXA₂ and, in consequence, produce the observed hemodynamic alterations.

Our findings, however, do not support this hypothesis. In the present experimental situation infusion of prostacyclin did not influence blood flow or flow regulation during delayed postischemic hyperperfusion following one hour of cerebrocirculatory arrest. A methodological error due to inactivation of PGI₂ before or during infusion can be excluded because its vasodilatory effect was confirmed by the fall of systemic arterial pressure,6 and its effect on platelet function by the reduction of platelet aggregability.9-11 It is not likely, either, that the drug did not reach the brain vessels because previous studies have shown that in animals with functional recovery blood reperfusion is not disturbed.3 However, it should be considered that the absence of cerebral vascular effects may be due to a reduced activity of PGI₂ at the effector site in the vascular smooth muscle cells, e.g. by a disturbed passage across the cerebrovascular wall.

As has been shown in previous experiments, disseminated intravascular coagulation develops during the ischemic impact being initiated by an activation of factor XII and increased platelet aggregation in the
microvasculature. The activation of blood platelets as well as of the procoagulant and fibrinolytic system with liberation of 5-hydroxytryptamine and other proaggregatory compounds from the platelets and formation of fibrin degradation products has been shown to increase the toxicity of cerebral and other vessels. The failing response of PGI2 on the vascular smooth muscles could, therefore, also be influenced by this disturbance of the haemostatic system which was not affected by the PGI2 infusion started 2 hours postischemia.

Our findings, on the first sight, seem to contradict the earlier observations of Hallenbeck and Furlow who reported an amelioration of post-ischemic recirculation by prostacyclin. However, these authors studied the early recirculation phase and not the delayed postischemic hypoperfusion phenomenon. Immediately after ischemia, recirculation may be impaired by the so-called no-reflow phenomenon which is the combined result of intravascular aggregation of blood particles, increased vascular tone, microvascular compression by swollen glial cells and post-ischemic hypotension. Prostacyclin seems to improve this disturbance by reducing platelet aggregation because the flow-promoting effect can be enhanced by additional application of indomethacin. Indomethacin, besides its effect on prostacyclin synthesis, also causes an inhibition of thromboxane A2 which is a platelet aggregator. Inhibition of these two compounds and substitution of prostacyclin, therefore, considerably decreases platelet aggregability.

In the present investigation, the no-reflow phenomenon was prevented in a different way. Since the degree of no-reflow is mainly determined by the degree of microvascular compression and the post-ischemic perfusion pressure, blood pressure was increased immediately before recirculation to hypertensive levels, and perivascular glial swelling was reduced by osmotherapy. With this approach it was possible to prevent the no-reflow phenomenon without application of specific drugs, and the effect of prostacyclin could be studied at a later stage without previous interference with prostaglandin metabolism. The absence of a hemodynamic effect of prostacyclin after 2 hours' recirculation, therefore, is a strong argument against its role for the development of post-ischemic hypoperfusion. However, if prostacyclin synthetase is inactivated by free radicals, as suggested by Siesjö, it is conceivable that not only the enzyme but also the prostacyclin effector site at the vascular smooth muscle is destroyed. Substitution of prostacyclin, in this case, would be without hemodynamic effects. Further studies, therefore, should focus on this particular problem.

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Intracranial arteriolar arterial pressure and retinal vascular changes in diseases and the distal segments of the bifurcation of the pathy of unknown origin and the site of occurrence is not clear. The disease is a chronic inflammatory arterio-occlusive thromboarteriopathy (Takayasu's disease) is RELATIONSHIP among cervical arterial stenoses, microaneurysms and the distal segments of the bifurcation of the.


Occlusive Thromboarteriopathy (Takayasu’s Disease): Cervical Arterial Stenoses, Retinal Arterial Pressure, Retinal Microaneurysms and Prognosis

KAICHIRO ISHIKAWA, M.D.,* MASANOBU UYAMA, M.D.,† AND KUNIO ASAYAMA, M.D.†

SUMMARY Eighty-one young Japanese patients with occlusive thromboarteriopathy (Takayasu’s disease) were classed into three groups according to the degree and extent of diameter stenosis in the 4 cervical arterial systems, as determined by serial aortography. Class I was made up of 63 patients with 70% or greater stenosis in less than 3 systems, including 33 patients without systemic hypertension (Class Ia). Class II was made up of 6 patients with 70% or greater stenosis in 3 systems and less than 50% stenosis in the remaining 1, including 5 patients without systemic hypertension (Class IIa). Class III was made up of 12 patients with 70% or greater stenosis in 3 systems and 50% or greater stenosis in 1 system. Ophthalmodynamometric systolic pressure in patients in Class III was significantly lower than in patients in Class IIa (p < 0.001), but there was no significant difference between patients in Classes Ia and IIa. Microaneurysms and/or arteriovenous anastomoses in the retinal vessels were found in all but one patient in Class III and in only one patient in combined Classes I and II. These results indicate that each of the ophthalmodynamometric values and fundoscopic findings are very helpful in identifying the markedly severe occlusive lesions (Class III) of the 4 cervical arterial systems. In this chronic disease, however, angiography is most useful for evaluation of these severe lesions, to monitor progression from Classes I and II to Class III, in which the prognosis is rather poor.

RELATIONSHIP among cervical arterial stenoses, retinal arterial pressure and retinal vascular changes in occlusive thromboarteriopathy (Takayasu’s disease) is not clear. The disease is a chronic inflammatory arteriopathy of unknown origin and the site of occurrence is the aorta and/or its main branches. Intracranial arteries and the distal segments of the bifurcation of the common carotid arteries and those of the vertebral arteries are usually spared in this disease. Therefore, these patients appropriately serve as unique subjects for assessment of the influence of gradually developing stenoses in the proximal segments of the aortic arch vessels, as related to retinal arterial pressure and ischemic retinopathy. A severe retinopathy of arteriovenous anastomoses and microaneurysms, a characteristic manifestation of Takayasu’s retinopathy is one of the ominous signs in prediction of a poor prognosis in patients with this disease. Consequently, it is very important to investigate stepwise, the cervical arterial stenoses which cause severe ischemic retinopathy.

From *The 3rd Division, Department of Internal Medicine and †Ophthalmology, Faculty of Medicine, Kyoto University, Kyoto, Japan. Address correspondence to: Dr. Kaichiro Ishikawa, The 3rd Division, Department of Internal Medicine, Faculty of Medicine, Kyoto University, Kyoto 606, Japan. Received October 26, 1982; revision accepted April 13, 1983.
No effect of prostacyclin on blood flow, regulation of blood flow and blood coagulation following global cerebral ischemia.

W van den Kerckhoff, K A Hossmann and V Hossmann

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