

Starting equations

$$\frac{D_{NO}}{r} \frac{d}{dr} \left( r \frac{dC_{NO}}{dr} \right) + R_{NO} = 0 \quad (1)$$

Simplify

$$D_{NO} \frac{d^2 C_{NO}}{dr^2} + \frac{D_{NO}}{r} \frac{dC_{NO}}{dr} = -R_{NO} \quad (2)$$

Compare to form

$$u'' + P(r)u' = F(r) \quad (3)$$

$$u = C_{NO}, \quad P(r) = \frac{1}{r}, \quad F(r) = -\frac{R_{NO}}{D_{NO}} \quad (4)$$

Taylor expansion

$$\begin{aligned} u_{i+1} &= u_i + u'_i \Delta r + \frac{1}{2} u''_i \Delta r^2, \\ u_{i-1} &= u_i - u'_i \Delta r + \frac{1}{2} u''_i \Delta r^2 \end{aligned}$$

Central difference

$$\begin{aligned} u'_i &= \frac{u_{i+1} - u_{i-1}}{2\Delta r}, \\ u''_i &= \frac{u_{i+1} - 2u_i + u_{i-1}}{\Delta r^2} \end{aligned}$$

Sub back into Eq. (3)

$$\frac{u_{i+1} - 2u_i + u_{i-1}}{h^2} + P_i \frac{u_{i+1} - u_{i-1}}{2h} = F_i, \quad h = \Delta r \quad (5)$$

where index  $i$  starts at 1.

Rearrange

$$\left(1 - \frac{h}{2} P_i\right) u_{i-1} + (-2)u_i + \left(1 + \frac{h}{2} P_i\right) u_{i+1} = h^2 F_i \quad (6)$$

Split into 5 sections. Solutions are in the general form  $A\tilde{\mathbf{u}} = B$ .

Let  $a_i = \frac{h}{2} P_i$ .

RBC core  $r_0 < r < r_1$

Governing equation:

$$\begin{aligned} \frac{D_{NO}}{r} \frac{d}{dr} \left( r \frac{dC_{NO}}{dr} \right) - \lambda_{core} C_{NO} &= 0 \\ F_i &= \frac{\lambda_{core} u_i}{D_{NO}} \end{aligned}$$

Boundary condition:

$$\begin{aligned} u'(0) &= 0 \rightarrow u_1 = u_2, \\ u'_{RBC}(r_1) &= \sigma, \quad \sigma = u'_{CFL}(r_1) \end{aligned}$$

Using second-order accurate one-sided difference approximation,  $j$  represents indexes in CFL domain.

$$\begin{aligned} \sigma &= \frac{-3u_j + 4u_{j+1} - u_{j+2}}{2h}, \\ \frac{3u_i - 4u_{i-1} + u_{i-2}}{2h} &= \sigma, \\ 3u_i - 4u_{i-1} + u_{i-2} &= 2h\sigma \end{aligned}$$

$$A = \begin{bmatrix} -2 & 1+a_2 & 0 & \cdots & \cdots & \cdots & \cdots & 0 \\ 1-a_3 & -2 & 1+a_3 & \ddots & & & & \vdots \\ 0 & 1-a_4 & -2 & \ddots & \ddots & & & \vdots \\ \vdots & \ddots & \ddots & \ddots & \ddots & & & \vdots \\ \vdots & & \ddots & \ddots & -2 & 1+a_{n-3} & 0 & 0 \\ \vdots & & & \ddots & 1-a_{n-2} & -2 & 1+a_{n-2} & 0 \\ \vdots & \cdots & \cdots & \cdots & 0 & 1-a_{n-1} & -2 & 1+a_{n-1} \\ 0 & \cdots & \cdots & \cdots & \cdots & 1 & -4 & 3 \end{bmatrix}, \tilde{\mathbf{u}} = \begin{bmatrix} u_2 \\ u_3 \\ \vdots \\ u_i \\ \vdots \\ u_n \end{bmatrix}, B = \begin{bmatrix} h^2 F_2 - (1-a_2)u_1 \\ h^2 F_3 \\ \vdots \\ h^2 F_i \\ \vdots \\ h^2 F_{n-1} \\ 2h\sigma \end{bmatrix} \quad (7)$$

CFL  $r_1 < r < r_2$   
Governing equation

$$\frac{D_{NO}}{r} \frac{d}{dr} \left( r \frac{dC_{NO}}{dr} \right) = 0$$

$$F_i = \frac{0}{D_{NO}}$$

Boundary condition:

$$u_{CFL}(r_1) = u_{RBC}(r_1),$$

$$u'_{CFL}(r_2) = \phi, \phi = u'_{EC}(r_2),$$

Using one-sided difference approximation again,  $i$  is CFL,  $j$  is RBC.

$$\phi = \frac{-3u_k + 4u_{k+1} - u_{k+2}}{2h},$$

$$\frac{3u_i - 4u_{i-1} + u_{i-2}}{2h} = \phi,$$

$$3u_i - 4u_{i-1} + u_{i-2} = 2h\phi$$

$$A = \begin{bmatrix} -2 & 1+a_2 & 0 & & & & & 0 \\ 1-a_3 & -2 & 1+a_3 & \ddots & & & & \vdots \\ 0 & 1-a_4 & -2 & \ddots & \ddots & & & \vdots \\ \vdots & \ddots & \ddots & \ddots & \ddots & & & \vdots \\ \vdots & & \ddots & \ddots & -2 & 1+a_{n-3} & 0 & 0 \\ \vdots & & & \ddots & 1-a_{n-2} & -2 & 1+a_{n-2} & 0 \\ \vdots & \cdots & \cdots & \cdots & 0 & 1-a_{n-1} & -2 & 1+a_{n-1} \\ 0 & \cdots & \cdots & \cdots & \cdots & 1 & -4 & 3 \end{bmatrix}, \tilde{\mathbf{u}} = \begin{bmatrix} u_2 \\ u_3 \\ \vdots \\ u_i \\ \vdots \\ u_n \end{bmatrix}, B = \begin{bmatrix} h^2 F_2 - (1-a_2)u_1 \\ h^2 F_3 \\ \vdots \\ h^2 F_i \\ \vdots \\ h^2 F_{n-1} \\ 2h\phi \end{bmatrix} \quad (8)$$

EC  $r_2 < r < r_3$  is same method as CFL, only difference is in the  $R_{NO}$  term.  
Governing equation

$$\frac{D_{NO}}{r} \frac{d}{dr} \left( r \frac{dC_{NO}}{dr} \right) + R_{NO} = 0$$

$$F_i = -\frac{R_{NO_{max}}}{D_{NO}} \frac{P_{O_2}}{P_{O_2} + K_{m,eNOS}}$$

Boundary condition:

$$\begin{aligned} u_{EC}(r_2) &= u_{CFL}(r_2), \\ u'_{EC}(r_3) &= \gamma, \quad \gamma = u'_{VW}(r_3) \end{aligned}$$

VW  $r_3 < r < r_4$  is same method as CFL, only difference is in the  $R_{O_2}$  term.

Governing equation

$$\begin{aligned} \frac{D_{NO}}{r} \frac{d}{dr} \left( r \frac{dC_{NO}}{dr} \right) - \lambda_{vw,t} C_{NO} &= 0 \\ F_i &= \frac{\lambda_{vw,t} u_i}{D_{NO}} \end{aligned}$$

Boundary condition:

$$\begin{aligned} u_{VW}(r_3) &= u_{EC}(r_3), \\ u'_{VW}(r_4) &= \delta, \quad \delta = u'_T(r_4) \end{aligned}$$

T  $r_4 < r < r_5$ . Combined since same governing equation. Governing equation

$$\begin{aligned} \frac{D_{NO}}{r} \frac{d}{dr} \left( r \frac{dC_{NO}}{dr} \right) - \lambda_{vw,t} C_{NO} &= 0 \\ F_i &= \frac{\lambda_{vw,t} u_i}{D_{NO}} \end{aligned}$$

$$\begin{aligned} u_T(r_4) &= u_{VW}(r_4), \\ u'(r_5) &= 0 \rightarrow u_n = u_{n-1} \end{aligned}$$

$$A = \begin{bmatrix} -2 & 1+a_2 & 0 & \cdots & \cdots & \cdots & \vdots \\ 1-a_3 & -2 & 1+a_3 & \ddots & & & \vdots \\ 0 & 1-a_4 & -2 & \ddots & \ddots & & \vdots \\ \vdots & \ddots & \ddots & \ddots & \ddots & & \vdots \\ \vdots & & \ddots & \ddots & -2 & 1+a_{n-3} & 0 \\ \vdots & & & \ddots & 1-a_{n-2} & -2 & 1+a_{n-2} \\ 0 & \cdots & \cdots & \cdots & 0 & 1-a_{n-1} & -2 \end{bmatrix}, \quad \tilde{\mathbf{u}} = \begin{bmatrix} u_2 \\ u_3 \\ \vdots \\ u_i \\ \vdots \\ u_{n-1} \end{bmatrix}, \quad B = \begin{bmatrix} h^2 F_2 - (1-a_2)u_1 \\ h^2 F_3 \\ \vdots \\ h^2 F_i \\ \vdots \\ h^2 F_{n-1} - (1+a_{n-1})u_n \end{bmatrix} \quad (9)$$

$O_2$  is solved using the same matrix as  $NO$  except in RBC where it has a constant partial pressure.

RBC core  $r_0 < r < r_1$

$$P_{O_2} = 70$$

CFL  $r_1 < r < r_2$

Governing equation

$$\begin{aligned} \alpha \frac{D_{O_2}}{r} \frac{d}{dr} \left( r \frac{dP_{O_2}}{dr} \right) &= 0 \\ F_i &= \frac{0}{\alpha D_{O_2}} \end{aligned}$$

Boundary condition:

$$\begin{aligned} v_{CFL}(r_1) &= v_{RBC}(r_1), \\ v'_{CFL}(r_2) &= \phi, \quad \phi = v'_{EC}(r_2) \end{aligned}$$

EC  $r_2 < r < r_3$

Governing equation

$$\alpha \frac{D_{O_2}}{r} \frac{d}{dr} \left( r \frac{dP_{O_2}}{dr} \right) - R_{NO} = 0$$

$$F_i = \frac{R_{NO_{max}}}{\alpha D_{O_2}} \frac{P_{O_2}}{P_{O_2} + K_{m,eNOS}}$$

Boundary condition:

$$\begin{aligned} v_{EC}(r_2) &= v_{CFL}(r_2), \\ v'_{EC}(r_3) &= \gamma, \gamma = v'_{VW}(r_3) \end{aligned}$$

VW  $r_3 < r < r_4$

Governing equation

$$\begin{aligned} \alpha \frac{D_{O_2}}{r} \frac{d}{dr} \left( r \frac{dP_{O_2}}{dr} \right) - Q_{O_2 \max VW} \frac{P_{O_2}}{P_{O_2} + appK_m} &= 0 \\ appK_m &= K_m \left( 1 + \frac{C_{NO}}{C_{ref}} \right) \\ F_i &= \frac{Q_{O_2 \max VW}}{\alpha D_{O_2}} \frac{P_{O_2}}{P_{O_2} + K_m \left( 1 + \frac{u_i}{C_{ref}} \right)} \end{aligned} \tag{10}$$

Boundary condition:

$$\begin{aligned} v_{VW}(r_3) &= v_{EC}(r_3), \\ v'_{VW}(r_4) &= \delta, \delta = v'_T(r_4) \end{aligned}$$

T  $r_4 < r < r_5$

Governing equation

$$\begin{aligned} \alpha \frac{D_{O_2}}{r} \frac{d}{dr} \left( r \frac{dP_{O_2}}{dr} \right) - Q_{O_2 \max T} \frac{P_{O_2}}{P_{O_2} + appK_m} &= 0 \\ appK_m &= K_m \left( 1 + \frac{C_{NO}}{C_{ref}} \right) \\ F_i &= \frac{Q_{O_2 \max T}}{\alpha D_{O_2}} \frac{P_{O_2}}{P_{O_2} + K_m \left( 1 + \frac{u_i}{C_{ref}} \right)} \end{aligned} \tag{11}$$

Boundary condition:

$$\begin{aligned} v_T(r_4) &= v_{VW}(r_4), \\ v'_T(r_5) &= 0 \rightarrow v_n = v_{n-1} \end{aligned}$$