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# Parameter Optimisation for *FCcRIy* Pathway to Two Different Datasets Using Least-Squares Optimisation

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## ABSTRACT

Syk is a tyrosine kinase important to bridge the receptor ligation and downstream signallings such as  $Ca^{2+}$  and PI3K. Once the cell receptor binds with the ligand, FC $\epsilon$ RI $\gamma$  (ITAM receptor) is recruited and phosphorylated by Lyn. The phosphorylated ITAM then recruits protein tyrosine kinase (Syk). The previously developed FC $\epsilon$ RI $\gamma$  (FC $\epsilon$ ) model contained a greater level of complexity. This study aims to build a simple model of signalling of FC $\epsilon$  that still represents biological understanding. The parameter estimation is addressed using least-squares optimisation, which implements the Levenburg-Marquardt gradient method (greedy algorithm) to minimise an objective function. More importantly, this model was fitted to two data sets that captured a temporal FC $\epsilon$ , Syk and Grb2 phosphorylation. Model uncertainty often has done as an analysis that is carried out after model construction and calibration have been completed. This study assessed for sensitivity to parameter choices and model uncertainty to perform the analysis. The modular design principles are applied to the construction of the model. The model is designed to be reproducible. In other words, the model can be effectively applied in simulation conditions or optimised to new datasets for new experimental situations.

Keywords: FC cRIy pathway, mathematical model, modular, parameter optimisation, reproducible

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## INTRODUCTION

The high-affinity IgE receptor (FC $\epsilon$ RI $\gamma$ ) is a highly important receptor found in many immune cells, including natural killer cells, mast cells, monocytes, eosinophils, and dendritic cells (Maurer, et al., 1996; Lantz, et al., 1997). FC $\epsilon$ RI $\gamma$  (FC $\epsilon$ ) involves determining immune cells' response toward target cells by secreting cytokines (Benhamou et al., 2019). For years, many *in vivo* and *in silico* studies have been done to understand the FC $\epsilon$  downstream signalling pathway once the receptor has been activated (Faeder et al., 2003; Lantz et al., 1997; Maurer et al., 1996; Tsang et al., 2008; Blank et al., 2021; Gregorio, 2019; Benhamou et al., 2019).

Activation of FC $\epsilon$  by Lyn-catalysed phosphorylation recruits Syk. Syk binding to FC $\epsilon$  is activated via phosphorylation by Lyn. Phosphorylated Syk then phosphorylates GRB2associated binding protein 2 (GAB2), which leads to activation of the phosphoinositide 3-kinase (PI3K) signalling cascade. FC $\epsilon$  exists in tetrameric form, which has an  $\alpha$ -chain, a  $\beta$ -chain, and two disulfide-linked  $\gamma$ -chain (Faeder et al., 2003). The  $\alpha$  subunit can be bound to a ligand or unbound, a  $\beta$ -chain has four possible states of phosphorylation, and two disulfidelinked  $\gamma$ -chain has six states of phosphorylations.

Previously Faeder et al. (2003) developed a model of early signalling of  $FC\epsilon$ . However, this model contained a greater level of complexity. In total, there are 300 possible states of receptor subunits for FC $\epsilon$  (Faeder et al., 2003). Faeder's model focuses on the early events in FC $\epsilon$  signalling, consisting of 3 components, 15 reaction classes and 21 rate constants. The parameters in Faeder's model were modified to fit the model to experimental data.

In this model, this study aims to build a simple model of signalling of FC $\epsilon$  that still represents biological understanding and can fit different experimental data sets. The idea is to develop a simple model with fewer unknown parameters that need to be predicted during optimisation. Further, the uncertainty in the model can be reduced. The model was built in component that can be mobilised and reusable to be included in larger pathway model. As described in Faeder et al. (2003), the early events in FC $\epsilon$  signalling can be described as a linear sequence. Here, a model that consists of 4 components, 7 reaction classes and 7 rate constants is developed. The model was optimised using the Levenburg-Marquardt gradient method (greedy algorithm) to minimise an objective function described later in Methodology.

This study assumed that FC $\epsilon$  is in an aggregated state and ready for binding

with other reactants to avoid complexity. First, the activated FC $\epsilon$  receptor is transphosphorylated by pLyn, available at the cell surface. Then, the recruitment of Syk follows it to the membrane surface, which is subsequently transautophosporylated by phosphorylated FC $\epsilon$ . Finally, inactive Grb2 binds to phosphorylated Syk and becomes phosphorylated (Figure 1).

Previous studies have shown that ITAM phosphorylation increases Syk activity and modulates Syk potency (Tsang et al., 2008). It must be noted that concentration



*Figure 1*. Schematic diagram of the FCcRIγ signalling pathway

above 10  $\mu$ M ITAM can inhibit Syk activity (Tsang et al., 2008). Phosphorylated Syk then phosphorylates Grb2 (growth-factor-receptor-bound protein 2). Phosphorylation of Grb2 leads to the activation of PI3K downstream signalling. In order to match the available experimental data, this study assumed that the Lyn kinase is rephosphorylated for recruitment of additional pLyn into the system to ensure that the system has enough supply of phosphorylated Lyn. The equation associated with this reaction is in Equation 1

$$Lyn + Pi \stackrel{\kappa_3}{\to} pA = \pi r^2 Lyn \tag{1}$$

where Pi denotes a phosphate, k denotes a forward reaction, and pLyn denotes a phosphorylated Lyn.

The flux associated with this reaction is in Equation 2

$$J3 = k_3[Pi][Lyn]$$
<sup>[2]</sup>

where J denotes a flux, k denotes kinetic constant, and P<sub>i</sub> denotes a phosphate.

The model was fitted to two published sets of experimental data. One was published by Tsang et al. (2008) and the other by Faeder et al. (2003). Experimental detail on Syk activity in B cells is presented in the study by Tsang et al. (2008), which captures the molecular mechanism of Syk activation in vitro. The studies explore Syk activity under various conditions: activities of phosphorylated and non-phosphorylated Syk; activation of Syk by different receptors; and whether Syk is an OR-gate type of molecular switch. Syk activity is required for more than one hour to induce activation of downstream reactions, so the experiment was run for 3600 s. For that reason, the model simulation time was set for 3600 s. For the FC $\epsilon$  model, there are 10 unknown parameters to fit this data.

In one of the observations, Tsang looked at the activity of Syk when bound to FC $\epsilon$ . The experimental data fitted the model for temporal Grb2 phosphorylation by dephosphorylated Syk with 1  $\mu$ M FC $\epsilon$ . The Syk concentration used in the experiment was 0.005  $\mu$ M. Experimental data points were digitised from Supplementary Data 2 in Tsang et al. (2008). The observed experimental data points were from spectrophotometrical measurements of phosphorylation in a single representative experiment (Tsang et al., 2008). The binding of Syk to the receptor eliminated the lag phase in Grb2 phosphorylation.

The second data set was adapted from Faeder et al. (2003). In their simulation, Faeder et al. (2003) looked at the pathway in rat basophilic leukaemia (RBL) cells. The cell density and volume were assumed to be  $1 \times 10^6$  cells/ml and  $1.4 \times 10^9$  ml. The observed experimental data points were from densitometric phosphorylation measurements in a single representative experiment (Faeder et al., 2003; Wofsy et al., 1995). In addition, experimental data points were digitised from Faeder et al. (2003). The setup of the two experiments is listed in Table 1.

Item	Tsang et al. (2008)	Faeder et al. (2003)
Cell type	B cell	RBL cell (rat basophilic cell)
Ligand	nil	IgE dimer
Receptor	$FC\epsilon RI\gamma$	$FC\epsilon RI\gamma$
Experiment temperature	37°C	27°C
Simulation time	3600 s	4200 s
Measurement method	Spectrophotometer	Flourescence plate
Concentration of Syk	0.005 µM	0.403 μΜ
Concentration of $FC \in RI\gamma$	1 µM	0.474 μM
Concentration of Lyn	nil	0.0332 μM

Table 1Comparison of experimental setting between Tsang et al. (2008) and Faeder et al. (2003)

The signalling pathway was built in CellML components (Cooling et al., 2008; Cooling et al., 2016), where component/s is built up using equations and variables related to a specific reaction. A good reproducible model needs a proper protocol for its users. Documentation of the equations, model optimisation to experimental data, and validation against other experimental data are not enough to make the model able to be reproduced (Waltemath et al., 2011). In such a protocol, the simulation conditions for a model are recorded, including how the model is used to simulate certain experiments and how the model generates a published output. Through its cross-platform modelling environment, OpenCOR, CellML, provides this facility where users can generate Simulation Experiment Description Markup Language (SED-ML) from the CellML model that they have built. SED-ML maintains the simulation protocol and records the simulation output (Waltemath et al., 2011). Users can easily know the simulation protocols such as the underlying mathematical equations, parameters used, type of solver algorithm, simulation time, and time steps taken for a particular experiment from a SED-ML format. When SED-ML is provided, other users can reference a model and publish figures accurately (Cooling et al., 2016). By using SED-ML, the model can be mobilised and re-used in other pathways that consist of the  $FC \in RI\gamma$  pathway.

## **MATERIALS AND METHODS**

1-

The cell's signalling pathways are represented using ordinary differential equations (ODEs). An ODE describes the concentration changes of a compound over time. In this model, flux is defined,  $J_1$  for each reaction in the comprehensive pathway, which helps to simplify the description of the equations. The flux typically comprises the concentration of species (denoted by [...] throughout this study) and rate constants  $k_f$  (forward reaction) and  $k_r$  (reverse reaction) in the law of mass action.

The reactions associated with this model are as in Equations 3-6

1.

$$FC\varepsilon + pLyn \stackrel{\kappa_1}{\leftrightarrow} pFC\varepsilon Lyn \stackrel{\kappa_2}{\to} pFC\varepsilon + Lyn,$$
<sup>[3]</sup>

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$$Lyn + Pi \xrightarrow{k_3} pLyn, \tag{4}$$

$$pFC\varepsilon + Syk \stackrel{k_4}{\leftrightarrow} pFC\varepsilon Syk \stackrel{k_5}{\to} FC\varepsilon + pSyk,$$
[5]

$$pSyk + Grb2 \stackrel{k_6}{\leftrightarrow} pSykGrb2 \stackrel{k_7}{\rightarrow} Syk + pGrb2.$$
 [6]

The fluxes associated with these reactions are as in Equations 7-13

$$J1 = k_1[FC\varepsilon][pLyn] - k_{-1}[pFC\varepsilon Lyn],$$
[7]

$$J2 = k_2 [pFC \varepsilon Lyn],$$
<sup>[8]</sup>

$$J3 = k_3[Pi][Lyn], [9]$$

$$J4 = k_4[pFC\varepsilon][Syk] - k_{-4}[pFC\varepsilon Syk],$$
<sup>[10]</sup>

$$J5 = k_5[pFC\varepsilon Syk],$$
[11]

$$J6 = k_6 [pSyk] [Grb2] - k_{-6} [pSykGrb2],$$
[12]

$$J7 = k_7 [pSykGrb2].$$
<sup>[13]</sup>

The ODEs associated with these reactions are as in Equations 14-24

$$\frac{pFC}{dt} = -J1 + J5,$$
[14]

$$\frac{pLyn}{dt} = -J1 + J3,$$
[15]

$$\frac{pFCLyn}{dt} = J1 - J2,$$
[16]

$$\frac{pFC}{dt} = J2 - J4,$$
[17]

$$\frac{pLyn}{dt} = J2 - J3,$$
[18]

$$\frac{pSyk}{dt} = -J4 + J7,$$
[19]

$$\frac{pFCSyk}{dt} = J4 - J5,$$
[20]

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$$\frac{pSyk}{dt} = J5 - J6,$$
[21]

$$\frac{Grb2}{dt} = -J6,$$
[22]

$$\frac{pSykGrb2}{dt} = J6 - J7,$$
[23]

$$\frac{pGrb2}{dt} = J7.$$
[24]

## **Model Optimisation and Sensitivity Analysis**

In this study, the model parameter optimisation of CellML models was done using Python inside the OpenCOR tool. The parameter estimation implements the Levenburg-Marquardt gradient method (greedy algorithm) to minimise an objective function (Moré, 1978). The first step is to define the objective function to minimise. This study, therefore, aims to randomly sample parameter space with suitable coverage to be able to assure that a solution can be found close to a global minimum (Saltelli et al., 2004). This study uses a sampling method to create a random sample of parameters within identified boundaries. The parameter samples were generated using the Saltelli sample function stored in Python modules.

In general, sensitivity analysis is the study of the changes of the optimal solution when there are changes in the constant parameters (Wallace, 2000). A sensitivity analysis was performed to determine the sensitivity of constant rate parameters in the model. Each of these parameters determines changes in the model. Therefore, each of these parameters determines changes in the model. This study initialised the model with the best fit values for each parameter and ran it multiple times over the parameter range. Each parameter was varied from its best fit value over a range of values. The error was defined as the difference between the model solution and experimental data values at each time point. The sum of the squared errors was used to quantify overall error over time (Sobol, 2001; Saltelli et al., 2010).

The sum of the squared errors was used to quantify the overall error over time. A dip in error shows the parameters sit at or very close to the minimum. However, the minimum can also be right at the edge of the parameter space. It means this parameter value needs to be around the 'minimum'.

## Model Parameters and Initial Conditions for Fitting to Experimental Data Set 1 [Tsang et al. (2008)]

This study first estimates B cell experimental data parameters from Tsang et al. (2008). Then, experimental data points were digitised from Supplementary Data 2 in Tsang et al. (2008).

The observed experimental data points were from spectrophotometrical measurements of phosphorylation in a single representative experiment (Tsang et al., 2008). The binding of Syk to the receptor eliminated the lag phase in Grb2 phosphorylation.

For the model, this study adapted the recruitment of additional Lyn to the phosphorylated ITAMs. The boundary conditions used in this fitting are between  $10^{-3}$  and  $10^2$  for the lower and upper bound (Cooling, 2007). However, the range expands when optimisation error potentially decreases with a larger boundary. The number of samples used for the fitting is 500, generating 12000 samples. The parameters to be fitted are  $k_{f1}$ ,  $k_{f2}$ ,  $k_{f3}$ ,  $k_{f4}$ ,  $k_{f5}$ ,  $k_{f6}$ ,  $k_{r7}$ ,  $k_{r4}$ , and  $k_{r6}$ .

The initial conditions adapted from the experiment are shown in Table 2. This study estimated the concentration of Grb2 and pLyn at 6.47  $\mu$ M and 6.5  $\mu$ M. The phosphorylated (except Lyn) and complex reactants are assumed to be 0 at time 0 s. The URL link for the Python script that runs fitting to Tsang data for this model, the sensitivity analysis code for the fitted parameter and the model it runs can be found in Supplementary Data 1.

Table 2 Initial conditions for the  $FC \in RI\gamma$  signalling model in parameter fitting to experimental data from Tsang et al. (2008)

Parameter	Value	Units	Source
FCε	1	μΜ	(Tsang et al., 2008)
Syk	0.005	μΜ	(Tsang et al., 2008)
Grb2	6.47	μΜ	Estimation
pLyn	6.5	μΜ	Estimation

## Model Parameters and Initial Conditions for Fitting to Experimental Data Set 2 [Faeder et al. (2003)]

The observed experimental data points were from densitometric phosphorylation measurements in a single representative experiment. Experimental data points were digitised from Faeder et al. (2003). Tsang et al. (2008) and Faeder et al. (2003) used different experimental protocols, so the initial conditions in this model representing these protocols differed. Some initial conditions were derived from the experimental protocol reported, and others were estimated during model fitting. In order to achieve the fit, several estimated initial conditions had to be changed from those estimated in Tsang's model. For fitting to Faeder's data, the concentration of FC $\epsilon$  is 0.0474  $\mu$ M, the concentration of Syk is 0.025 µM, and the concentration of pLyn is 0.0474 µM. These changes are listed in Table 3. Parameter fitting was then performed by fixing the previously fitted model's parameters. This study fixed the value of  $k_{12}$ ,  $k_{14}$ ,  $k_{16}$  and  $k_{17}$  (Table 4). In this subsequent fitting, for constant forward rate parameters kfl and reverse rate parameters  $k_{rl}$ ,  $k_{r4}$  and  $k_{r6}$ that showed distinct changes in model behaviour on one side of the minimum, this study varied the parameters only over the range that model predictions for the Tsang data were not sensitive to these parameters. The parameter that is going to fit and the boundaries are listed in Table 5. The URL link for the Python script that runs fitting to Faeder data for this

al. (2003)	,			
Parameter	Estimation 1	Source	Estimation 2	Units
$FC\epsilon$	0.474	(Faeder et al., 2003)	0.0474	$\mu M$

(Faeder et al., 2003)

nil

Initial conditions for the  $FC \in RV$  signalling model in parameter fitting to experimental data from Faeder et

pLyn 0.0332 (Faeder et al., 2003) 0.0474

model, the sensitivity analysis code for the fitted parameter and the model it runs can be found in Supplementary Data 1.

0.432

nil

## **Model Uncertainty**

Table 3

Syk

Grb2

Parameter optimisation sought a 'best fit' solution, that is, the set of parameters that provides the minimum error. However, this best fit may not be the only feasible model parameterisation closely similar to the experimental data. There may be alternate sets of parameters that provide solutions close to that data.

This study reconsiders the experimental data to determine whether any 'worse' fits still meet sensible criteria. This study searches for physiologically feasible parameter combinations to replicate the model function. To do that, this study search Table 4 Fixed kinetic parameters in the subsequence fitting

0.025

0.01

μM

μM

μM

Parameter	Value	Units
$k_{f2}$	0.0081	<b>s</b> <sup>-1</sup>
$k_{_{f4}}$	10.5797	$\mu M^{-1}.\mathrm{s}^{-1}$
$k_{f6}$	0.4143	$\mu M^{-1}.\mathrm{s}^{-1}$
$k_{f7}$	11.4185	$s^{-1}$

Table 5

Kinetic parameters to fit data in Faeder et al. (2003) and the boundaries condition

Parameter	Lower bound	Upper bound
$k_{f1}$	10-1.5	10 <sup>2</sup>
$k_{f3}$	10-3	10 <sup>2</sup>
$k_{f5}$	$10^{1}$	10 <sup>2</sup>
$k_{r1}$	10-3	10 <sup>1</sup>
$k_{r4}$	10-3	$10^{0.5}$
$k_{r6}$	10-3	$10^{0}$
Pi	10-3	10 <sup>2</sup>

for other solutions within the parameter space that predicts concentrations of pGrb2, which is the end output of the pathway, that lie within  $0.01 \mu M$  but may not be the 'best fit' returned by an optimiser.

This study sample parameter space to cover the range of feasible parameter combinations without requiring an unfeasibly large number of model runs. This study achieves this by using the inbuilt Saltelli sampling function in the python SALib library. The Saltelli sampling function generates  $N \times (2D + 2)$  samples, where N is a user-defined number, and D is the number of unknown parameters (the dimension of the parameter space which is aim to sample (Saltelli, 2002). To adequately cover the parameter space, based on the parameter swipe of the model, the N was set at 100, resulting in 1800 simulations.

## **RESULTS AND DISCUSSION**

## Parameter Fitting to Data from Tsang et al. (2008) and Sensitivity Analysis

The repository associated with this model can be found at https://github.com/Nurulizza/ FCepsilonRI.git and the SED-ML for the fitted model can be found at https://github.com/ Nurulizza/FCepsilonRI/blob/master/FCepsilonRI\_Tsangdata.sedml. The simulated Grb2 phosphorylation was a good fit with the experimental observations of Tsang et al. (2008). The model and observed experimental kinetics of phosphorylation of Grb2 are shown in Figure 2. The root mean square (RMS) error for the model is 0.0695  $\mu$ M. The RMSE is 1.07% of the peak concentration. Visual inspection of the graph confirms this. The best fit to this data set was achieved with the parameters listed in Table 6.



*Figure 2*. The best fit for phosphorylation of Grb2

*Note.* The best fit for phosphorylation of Grb2. Blue curves show model predictions, and red points are observed experimental data in Tsang et al. (2008). The SED-ML is available at https://github.com/Nurulizza/FCepsilonRI/blob/master/FCepsilonRI Tsangdata.sedml.

Table 6

Fitted parameters to experimental data from Tsang et al. (2008)

Parameter	Description	Value	Source
$k_{f1}$	FC-pLyn binding rate	58.3902 µM <sup>-1</sup> .s <sup>-1</sup>	Fitted value
$k_{f^2}$	FC phosphorylation rate	$0.00817889 \text{ s}^{-1}$	Fitted value
$k_{f3}$	Lyn phosphorylation rate	$1.0887 \ \mu M^{-1}.s^{-1}$	Fitted value
$k_{_{f4}}$	pFC-Syk binding rate	10.5797 μM <sup>-1</sup> .s <sup>-1</sup>	Fitted value
$k_{f5}$	Syk phosphorylation rate	63.6727 s <sup>-1</sup>	Fitted value
$k_{f6}$	pSyk-Grb2 binding rate	0.414288 µM <sup>-1</sup> .s <sup>-1</sup>	Fitted value
$k_{f7}$	Grb2 phosphorylation rate	11.4185 s <sup>-1</sup>	Fitted value
$k_{r1}$	FC-pLyn dissociation rate	0.013548 s <sup>-1</sup>	Fitted value
$k_{r4}$	pFC-Syk dissociation rate	0.0806961 s <sup>-1</sup>	Fitted value
<i>k</i> <sub><i>r</i>6</sub>	pSyk-Grb2 dissociation rate	0.731255 s <sup>-1</sup>	Fitted value

Knowledge of which parameters the model is sensitive to helped establish which parameters needed to be fixed when this study tried to predict another set of data (e.g. the phosphorylation of Grb2). Figure 3 present the differences in pSyk solutions within the feasible parameter range for all rate constants in this model. This study varied each parameter, one at a time, over the feasible parameter range to check how much they affected the solutions while keeping other rate constant parameters fixed.

For forwarding constants  $k_{j2}$ ,  $k_{j4}$ ,  $k_{j6}$ , and  $k_{j7}$ , there is a minimum error corresponding to the fitted parameter value, which showed the fitting had converged to something appropriate where most parameters were very close to at least a local minimum in the parameter space. It means the



*Figure 3.* The sensitivity of the FC $\epsilon$ RI $\gamma$  model solutions around the 'best fit' to a selection of parameters *Note.* The model was initialised with the best fit values for each parameter (fit value) and run multiple times over the parameter range as assumed (shown on the x-axis). Error is shown in red.

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*Figure 3* (continue). The sensitivity of the FC $\in$ RI $\gamma$  model solutions around the 'best fit' to a selection of parameters

*Note.* The model was initialised with the best fit values for each parameter (fit value) and run multiple times over the parameter range as assumed (shown on the x-axis). Error is shown in red.

model is sensitive to those parameters. The sensitive parameters are the dissociation rate of phosphorylated FC $\epsilon$  and Lyn complex,  $k_{f2}$ , an association of phosphorylated FC $\epsilon$  and Syk,  $k_{f4}$ , an association of pSyk and Grb2,  $k_{f6}$ , and dissociation of pSyk and Grb2 complex,  $k_{f7}$ . These parameters remained fixed in subsequent simulations. Constant forward rate parameters  $k_{f1}$  and reverse rate parameters  $k_{r1}$ ,  $k_{r4}$ , and  $k_{r6}$  showed distinct changes in model behaviour on one side of the minimum. In this case, these parameters can vary when fitting to another set of data in the range around the minimum the line is flat. A flat line was found for  $k_{f1}$  and Pi, showing that the model is not sensitive to those parameters (data not shown). Therefore, the latter two types of parameters could be varied over the whole defined range in fitting to subsequent data sets.

## Multi-Objective Parameter Fitting to Data Set from Faeder et al. (2003) and Sensitivity Analysis

The repository associated with this model can be found at https://github.com/Nurulizza/ FCepsilonRI.git and the SED-ML for the fitted model can be found at https://github.com/ Nurulizza/FCepsilonRI/blob/master/FCepsilonRI Faederdata.sedml. In this subsequent

parameter fitting, the concentration of FC $\epsilon$  is 0.0474  $\mu$ M, the concentration of Syk is 0.025  $\mu$ M, and the concentration of pLyn is 0.0474  $\mu$ M, as listed in Table 2. Parameter fitting was then performed by fixing parameters  $k_{f2}$ ,  $k_{f4}$   $k_{f6}$ , and  $k_{f7}$  (Figure 5 and Table 7). The list of kinetic rate constants and the boundaries are given in Tables 4 and 5. Finally, the best fit was achieved with the parameters listed in Table 6. The model and the observed experimental Faeder et al. (2003) kinetics of phosphorylation of FC and Syk are shown in Figure 4. The RMSEs are 0.0089  $\mu$ M for FC phosphorylation and 0.0022  $\mu$ M for Syk phosphorylation. These RMS errors are approximately 10% of the peak values for pFC and pSyk. Along with Figure 4, these best fits are not perfect representations of the data. However, the available data are relatively sparse in time and appear to contain more noise, so it may not be possible to obtain a better fit for these data points.



*Figure 4.* The best fit for phosphorylation of FCε (left) and phosphorylation of Syk (right) *Note.* Blue curves show model predictions, and red points are observed experimental data from Faeder et al. (2003). The SED-ML is available at https://github.com/Nurulizza/FCepsilonRI/blob/master/FCepsilonRI\_ Faederdata.sedml.

Table 7 The parameters for the  $FC \in RI\gamma$  model fitted to the data set from Tsang et al. (2008) and Faeder et al. (2003)

Parameter	Description	Value	Source	Status
$k_{f1}$	FC-pLyn binding rate	54.7678 μM <sup>-1</sup> .s <sup>-1</sup>	Fitted to Faeder et al. (2003)	Fit
$k_{f2}$	FC phosphorylation rate	$0.00817889 \ s^{-1}$	Fitted to Tsang et al. (2008)	Fix
$k_{f3}$	Lyn phosphorylation rate	$0.0035 \ \mu M^{-1}.s^{-1}$	Fitted to Faeder et al. (2003)	Fix
$k_{f4}$	pFC-Syk binding rate	10.5797 μM <sup>-1</sup> .s <sup>-1</sup>	Fitted to Tsang et al. (2008)	Fit
$k_{f5}$	Syk phosphorylation rate	33.7157 s <sup>-1</sup>	Fitted to Faeder et al. (2003)	Fit
$k_{f6}$	pSyk-Grb2 binding rate	$0.414288 \ \mu M^{-1}.s^{-1}$	Fitted to Tsang et al. (2008)	Fit
$k_{f7}$	Grb2 phosphorylation rate	11.4185 s <sup>-1</sup>	Fitted to Tsang et al. (2008)	Fix
$k_{r1}$	FC-pLyn dissociation rate	0.0031 s <sup>-1</sup>	Fitted to Faeder et al. (2003)	Fit
$k_{r4}$	pFC-Syk dissociation rate	0.1174 s <sup>-1</sup>	Fitted to Faeder et al. (2003)	Fit
$k_{r6}$	pSyk-Grb2 dissociation rate	0.481 s <sup>-1</sup>	Fitted to Faeder et al. (2003)	Fit

The fifth column listed parameters that can be fitted or fixed in the calibration of subsequent components due to the sensitivity analysis.



However, the model predictions shown in Figure 4 does not predict the experimentally observed drop in pSyk after 1000 s due to the formation of the pSyk-Grb2 complex. It may be because the concentration of Grb2 in the model was low (0.01  $\mu$ M), which limited its impact on pSyk. Therefore, the initial condition of Grb2 was fitted at 0.01  $\mu$ M to get a good fit to the experimental data. The sensitivity analysis of the model is shown in Figure 5.



*Figure 5.* The sensitivity of the FC $\in$ RI $\gamma$  model solutions around the 'best fit' to a selection of parameters *Note.* The sensitivity of the FC $\in$ RI $\gamma$  model solutions around the 'best fit' to a selection of parameters. The model was initialised with the best fit values for each parameter (fit value) and run multiple times over the parameter range as assumed (shown on the x-axis). Error is shown in red.

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The sensitivity analysis showed that the model fitted to the data measured by Faeder et al. (2003) is sensitive to the dissociation rate of phosphorylated FC $\epsilon$  and Lyn complex,  $k_{f2}$ , the phosphorylation rate of Lyn,  $k_{f3}$ , and Pi for the pFC. The same parameters, especially  $k_{f3}$  and Pi, however, do not contribute to pSyk variability. The model fitted the experimental data well with large values of parameters  $k_{f1}$ ,  $k_{f4}$ , and  $k_{f5}$ . Sensitivity analysis showed that the model converges toward minimum error for both FC and Syk phosphorylation on the right region of the parameter space. In contrast, parameters  $k_{r1}$  gave the best fit with small values when the model converges toward minimum error for FC and Syk phosphorylation on the left region of the parameter space. The pathway can be calibrated around the regions within the minimum error of parameters space. The model was not sensitive to the dissociation rate of pFC and Syk,  $k_{r4}$ , an association of pSyk and Grb2,  $k_{f6}$ , dissociation of pSyk and Grb2,  $k_{r6}$ , and phosphorylation of Subsequent model components that incorporate this pathway.

## **Model Uncertainty**

This custom script exported solution/s into a file within  $0.01\mu$ M pGrb2. After the script ran until the end of 1800 simulations, 228 alternate solutions were outputted to give different model behaviour predictions. The ssq of the alternate solutions range from 7.89e-31 to 0.0033. It means that over the range of parameter space assessed, the 'best fit' solution obtained from parameter fitting was not the only solution that provided physiological concentrations of cytokines. It means that the 'best fit' solution is not unique. It gives some confidence that the model provides many satisfactory solutions. The list of the alternate solutions is provided in Supplementary Data 3.

## CONCLUSION

This study developed and implemented a minimal and modular FC $\epsilon$  model in CellML. The results for phosphorylation of Grb2 for a data set from Tsang et al. (2008) were presented to validate and test for phosphorylation of FC $\epsilon$  and Syk in a data set from Faeder et al. (2003). In addition, a basic sensitivity analysis was performed to study the effect of the model constants' parameters on the model. The minimal and modular model allows modularity and increases the re-usability of the model in future work.

Lyn kinase phosphorylates  $FC\epsilon$  into phosphorylated FC. Re-phosphorylation of Lyn then occurs. Due to a lack of quantitative information about the aggregation rate of  $FC\epsilon$  in the literature, this study constructed a simple model of the system, in which the  $FC\epsilon$  was in the aggregation state. The model neglects aspects of the complex dynamics of the  $FC\epsilon$  system, which has previously been incorporated into mathematical descriptions. It means that although it captures the dynamics of the available experimental data adequately, it may be shown in the future to neglect important dynamics as more experimental data become available. The modular structure of this modelling approach would allow the component of the model to be replaced with a more detailed in the future, if necessary.

Phosphorylated FC $\epsilon$  activates Syk through binding activity. The activation of Syk eliminates the lag phase in Syk activity and results in a linear rate of Grb2 phosphorylation, indicating that FC $\epsilon$  binding activates Syk. Observations using the model displayed qualitatively similar pFC-Syk binding behaviour to the available experimental data.

Experimental work by Tsang et al. (2008) showed that Syk is an OR-gate switch, meaning that it can reach full activation either by ITAM binding or autophosphorylation activation. Tsang et al. (2008) demonstrated that activation by both stimuli works through the same mechanism, and the application of both stimuli is expected to give only a small increase in activity. The study's model supports this and implies that ITAM binding is sufficient to cause full activation of Syk. Phosphorylated Syk sustains activity over time to facilitate longer-term changes in cell signalling.

In this model, the Syk is an OR-gate switch, meaning that it can reach full activation with one factor by ITAM binding. The ability of Syk to reach full activation with a single stimulus helps to define the ability of Syk to sustain its activity over time, although after transient activation of ITAM to facilitate longer-term changes in cell signalling. For example, Syk activity is required for more than 1 hour to induce activation of NFAT transcription Tsang et al. (2008).

Phosphorylation of Grb2 is dependent on the available concentration of Syk and Grb2 in the cell. The simulation results showed that the model could produce quantitatively similar kinetic behaviour in the phosphorylation of Grb2 to the experimental data. Phosphorylation of Grb2 was initially linear with time and then plateaued after all the Grb2 was completely phosphorylated. The lag in the reaction time was eliminated.

The parameter space sampling could provide an alternate physiological solution. There are equally valid solutions based on the input data (large range of parameter bound) that give different predictions of the model behaviour. Considering the uncertainty of the model parameters prediction helps us understand more of the model behaviour. The alternate solution could be better than the 'best fit' models, although it is not 'optimal'. The alternate solution might help find systematic misfits to the data and constrain the range for each parameter value. It can be equally good predictions, which might be more robustly estimated.

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## **SUPPLEMENTARY DATA 1**

The python script that runs fitting to Tsang data for this model can be found at https://github.com/ Nurulizza/FCepsilonRI/blob/master/parameter\_ sweep fcepsilonri tsangdata.py

The model it runs is https://github.com/Nurulizza/ FCepsilonRI/blob/master/FCepsilonRI\_Tsangdata. cellml

The sensitivity analysis code for the fitted parameter can be found at https://github.com/Nurulizza/ FCepsilonRI/blob/master/sweep\_sensitivity\_ analysis\_fcepsilonri\_fittedtsang.py The python script that runs fitting to Faeder data for this model can be found at

https://github.com/Nurulizza/FCepsilonRI/blob/ master/parameter\_sweep\_fcepsilonri\_faederdata.py

The model it runs is

https://github.com/Nurulizza/FCepsilonRI/blob/ master/FCepsilonRI\_Faederdata.cellml

The sensitivity analysis code for the fitted parameter can be found at

https://github.com/Nurulizza/FCepsilonRI/blob/ master/sweep\_sensitivity\_analysis\_fcepsilonri\_ fittedfaeder.py

## **SUPPLEMENTARY DATA 2**

Code for model fitted to Tsang et al. (2008)

```
def model FCepsilonRI as
```

```
//
 *****
 //***
                             ***
        UNITS
 ******
 //
 def unit s as
   unit second;
 enddef:
 def unit uM as
   unit mole {pref: micro};
   unit liter {expo: -1};
 enddef;
 def unit per s as
   unit s {expo: -1};
 enddef:
 def unit uM_per_s as
   unit uM;
   unit s {expo: -1};
 enddef:
 def unit per uM per s as
   unit uM {expo: -1};
   unit s {expo: -1};
 enddef:
 //
```

```
*****
 //***
        COMPONENTS
                                   ***
 *****
 def comp environment as
   var t: s {pub: out};
 enddef;
 def comp FCepsilonRI as
   var t: s {pub: in};
   var J1: uM per s;
   var J2: uM per s;
   var J3: uM per s;
   var J4: uM per s;
   var J5: uM_per_s;
   var J6: uM_per_s;
   var J7: uM per s;
   var Lyn: uM {init: 0};
   var FC: uM {init: 1};
   var pFC: uM {init: 0};
   var Syk: uM {init: 0.005};
   var pSyk: uM {init: 0};
   var pGrb2: uM {init: 0};
   var pFCLyn: uM {init: 0};
   var pFCSyk: uM {init: 0};
   var pSykGrb2: uM {init: 0};
   var Pi: uM {init: 1.89585};
   var Grb2: uM {init: 6.47};
   var pLyn: uM {init: 6.5};
```

//var k\_fl: per\_uM\_per\_s {init: 58.3902}; //var k\_rl: per\_s {init: 0.013548};

```
//var k f2: per s {init: 0.00817889};
    //var k f3: per uM per s {init: 1.0887};
    //var k f4: per uM per s {init: 10.5797};
    //var k r4: per s {init: 0.0806961};
                                                       //
    //var k f5: per s {init: 63.6727};
    //var k f6: per uM per s {init: 0.414288};
                                                      def unit s as
    //var k r6: per s {init: 0.731255};
    //var k f7: per s {init: 11.4185};
                                                       enddef;
    var k_f1: per_uM_per_s {init: 58.69};
    var k r1: per s {init: 0.03};
    var k f2: per s {init: 0.00817889};
    var k f3: per uM per s {init: 1.0887};
                                                       enddef;
    var k f4: per uM per s {init: 4.6};
    var k_r4: per_s {init: 44.26};
    var k f5: per s {init: 10.89};
    var k f6: per uM per s {init: 19.70};
                                                       enddef;
    var k r6: per s {init: 0.0016};
    var k_f7: per_s {init: 29.53};
                                                         unit uM;
    J1 = k_f1*FC*pLyn-k_r1*pFCLyn;
    J2 = k f2*pFCLyn;
                                                       enddef;
    J3 = k_f3*Pi*Lyn;
    J4 = k_f4*pFC*Syk-k_r4*pFCSyk;
    J5 = k f5*pFCSyk;
    J6 = k f6*pSyk*Grb2-k r6*pSykGrb2;
    J7 = k f7*pSykGrb2;
                                                       enddef;
    ode(FC, t) = -J1+J5;
    ode(pFC, t) = J2-J4;
                                                       //
    ode(Syk, t) = -J4+J7;
    ode(pSyk, t) = J5-J6;
                                                      //***
    ode(Lyn, t) = J2-J3;
    ode(pLyn, t) = -J1+J3;
    ode(pFCLyn, t) = J1-J2;
                                                       //
    ode(pFCSyk, t) = J4-J5;
    ode(pSykGrb2, t) = J6-J7;
    ode(Grb2, t) = -J6;
    ode(pGrb2, t) = J7;
  enddef;
                                                       enddef;
 def map between environment and FCepsilonRI for
    vars t and t:
  enddef;
enddef;
Code for model fitted to Faeder et al. (2003)
def model FCepsilonRI as
 //
                                                         var Lyn: uM {init: 0};
 var FC: uM {init: 0.0474};
```

\*\*\*\*

```
//***
        UNITS
                               ***
 //*********************
*****
   unit second;
 def unit uM as
   unit mole {pref: micro};
   unit liter {expo: -1};
 def unit per_s as
   unit s {expo: -1};
 def unit uM_per_s as
   unit s {expo: -1};
 def unit per_uM_per_s as
   unit uM {expo: -1};
   unit s {expo: -1};
 *****
      COMPONENTS ***
 //****************
*****
 def comp environment as
   var t: s {pub: out};
 def comp FCepsilonRI as
   var t: s {pub: in};
   var J1: uM_per_s;
   var J2: uM per s;
   var J3: uM per s;
   var J4: uM per s;
   var J5: uM per s;
   var J6: uM per s;
   var J7: uM_per_s;
```

var pFC: uM {init: 0};

var Syk: uM {init: 0.025}; var pSyk: uM {init: 0}; var pGrb2: uM {init: 0}; var pFCLyn: uM {init: 0}; var pFCSyk: uM {init: 0}; var pSykGrb2: uM {init: 0}; var Pi: uM {init: 0.4027}; var Grb2: uM {init: 0.01}; var pLyn: uM {init: 0.0474}; var k\_f1: per\_uM\_per\_s {init: 54.7678}; var k r1: per s {init: 0.0031}; var k f2: per s {init: 0.00817889}; var k f3: per uM per s {init: 0.0035}; var k f4: per uM per s {init: 10.5797}; var k\_r4: per\_s {init: 0.1174}; var k f5: per s {init: 33.7157}; var k f6: per uM per s {init: 0.4143}; var k r6: per s {init: 0.481}; var k\_f7: per\_s {init: 11.4185};

J4 = k f4\*pFC\*Syk-k r4\*pFCSyk; J5 = k f5\*pFCSyk;J6 = k f6\*pSyk\*Grb2-k r6\*pSykGrb2; J7 = k f7\*pSykGrb2;ode(FC, t) = -J1+J5; ode(pFC, t) = J2-J4; ode(Syk, t) = -J4+J7; ode(pSyk, t) = J5-J6;ode(Lyn, t) = J2-J3; ode(pLyn, t) = -J1+J3; ode(pFCLyn, t) = J1-J2; ode(pFCSyk, t) = J4-J5;ode(pSykGrb2, t) = J6-J7;ode(Grb2, t) = -J6;ode(pGrb2, t) = J7;enddef;

def map between environment and FCepsilonRI for vars t and t; enddef;

1.FCepsilonRI/k f1: 27.5076

## **SUPPLEMENTARY DATA 3**

J1 = k f1\*FC\*pLyn-k r1\*pFCLyn;

FCepsilonRI/k\_f4: 22.5946 FCepsilonRI/k\_f5: 2.18697 FCepsilonRI/k\_f6: 5.94859 FCepsilonRI/k\_f7: 2.02145 FCepsilonRI/k\_r1: 0.0585569 FCepsilonRI/k\_r4: 1.6511 FCepsilonRI/k\_r6: 22.4174 Ssq[0.00086292 0.00086292]

 $J2 = k_f2*pFCLyn;$  $J3 = k_f3*Pi*Lyn;$ 

2.FCepsilonRI/k\_f1: 1.00451 FCepsilonRI/k\_f4: 24.8324 FCepsilonRI/k\_f5: 2.18697 FCepsilonRI/k\_f5: 5.94859 FCepsilonRI/k\_f7: 2.02145 FCepsilonRI/k\_r1: 0.0585569 FCepsilonRI/k\_r4: 1.6511 FCepsilonRI/k\_r6: 22.4174 Ssq[0.00080923] 0.00080923]

3.FCepsilonRI/k\_f1: 1.00451 FCepsilonRI/k\_f4: 22.5946 FCepsilonRI/k\_f5: 2.18697 FCepsilonRI/k\_f5: 5.94859 FCepsilonRI/k\_f7: 7.96833 FCepsilonRI/k\_r1: 0.0585569 FCepsilonRI/k\_r4: 1.6511 FCepsilonRI/k\_r6: 22.4174 Ssq[2.37725953e-152.37725953e-15]

4.FCepsilonRI/k\_f1: 1.00451 FCepsilonRI/k\_f4: 22.5946 FCepsilonRI/k\_f5: 2.18697 FCepsilonRI/k\_f6: 5.94859 FCepsilonRI/k\_f7: 2.02145 FCepsilonRI/k\_r1: 0.00652309 FCepsilonRI/k\_r4: 1.6511 FCepsilonRI/k\_r6: 22.4174 Ssq[0.00086435 0.00086435]

5.FCepsilonRI/k\_f1: 1.00451 FCepsilonRI/k\_f4: 22.5946 FCepsilonRI/k\_f5: 2.18697 FCepsilonRI/k\_f6: 5.94859 FCepsilonRI/k\_f7: 2.02145 FCepsilonRI/k\_r1: 0.0585569 FCepsilonRI/k\_r4: 0.536404 FCepsilonRI/k\_r6: 22.4174 Ssq[0.00068929 0.00068929]

6.FCepsilonRI/k\_f1: 1.00451 FCepsilonRI/k\_f4: 22.5946 FCepsilonRI/k\_f5: 2.18697 FCepsilonRI/k\_f6: 5.94859 FCepsilonRI/k\_f7: 2.02145 FCepsilonRI/k\_r1: 0.0585569 FCepsilonRI/k\_r4: 1.6511 FCepsilonRI/k\_r6: 0.0993277 Ssq[6.5012958e-21 6.5012958e-21]

7.FCepsilonRI/k\_f1: 1.00451 FCepsilonRI/k\_f4: 22.5946 FCepsilonRI/k\_f5: 2.18697 FCepsilonRI/k\_f5: 5.94859 FCepsilonRI/k\_f7: 2.02145 FCepsilonRI/k\_r1: 0.0585569 FCepsilonRI/k\_r4: 1.6511 FCepsilonRI/k\_r6: 22.4174 Ssq[0.0008684 0.0008684]

8.FCepsilonRI/k\_f1: 86.9868 FCepsilonRI/k\_f4: 10.4718 FCepsilonRI/k\_f5: 11.8637 FCepsilonRI/k\_f6: 43.0318 FCepsilonRI/k\_f7: 0.448092 FCepsilonRI/k\_r1: 0.00206278 FCepsilonRI/k\_r4: 0.0301642 FCepsilonRI/k\_r6: 0.00558561 Ssq[2.01998882e-22 2.01998882e-22] 9.FCepsilonRI/k\_f1: 3.17653 FCepsilonRI/k\_f4: 10.4718 FCepsilonRI/k\_f5: 11.8637 FCepsilonRI/k\_f6: 43.0318 FCepsilonRI/k\_f7: 0.448092 FCepsilonRI/k\_r1: 0.00206278 FCepsilonRI/k\_r4: 0.0301642 FCepsilonRI/k\_r6: 0.00558561 Ssq[2.90032855e-21 2.90032855e-21]

10.FCepsilonRI/k\_f1: 86.9868 FCepsilonRI/k\_f4: 53.5801 FCepsilonRI/k\_f5: 11.8637 FCepsilonRI/k\_f6: 43.0318 FCepsilonRI/k\_f7: 0.448092 FCepsilonRI/k\_r1: 0.00206278 FCepsilonRI/k\_r4: 0.0301642 FCepsilonRI/k\_r6: 0.00558561 Ssq[4.31725751e-19 4.31725751e-19]

11.FCepsilonRI/k\_f1: 86.9868 FCepsilonRI/k\_f4: 10.4718 FCepsilonRI/k\_f5: 21.8697 FCepsilonRI/k\_f6: 43.0318 FCepsilonRI/k\_f7: 0.448092 FCepsilonRI/k\_r1: 0.00206278 FCepsilonRI/k\_r4: 0.0301642 FCepsilonRI/k\_r6: 0.00558561 Ssq[3.46636391e-20 3.46636391e-20]

12.FCepsilonRI/k\_f1: 86.9868 FCepsilonRI/k\_f4: 10.4718 FCepsilonRI/k\_f5: 11.8637 FCepsilonRI/k\_f5: 43.0318 FCepsilonRI/k\_f7: 0.448092 FCepsilonRI/k\_r1: 0.00185173 FCepsilonRI/k\_r4: 0.0301642 FCepsilonRI/k\_r6: 0.00558561 Ssq[1.88627345e-17 1.88627345e-17]

13.FCepsilonRI/k\_f1: 86.9868 FCepsilonRI/k\_f4: 10.4718 FCepsilonRI/k\_f5: 11.8637 FCepsilonRI/k\_f6: 43.0318 FCepsilonRI/k\_f7: 0.448092 FCepsilonRI/k\_r1: 0.00206278 FCepsilonRI/k\_r4: 29.3612 FCepsilonRI/k\_r6: 0.00558561 Ssq[0.00065352 0.00065352]

14.FCepsilonRI/k\_f1: 86.9868 FCepsilonRI/k\_f4: 10.4718 FCepsilonRI/k\_f5: 11.8637 FCepsilonRI/k\_f5: 11.8637 FCepsilonRI/k\_f6: 43.0318 FCepsilonRI/k\_f7: 0.448092 FCepsilonRI/k\_r1: 0.00206278 FCepsilonRI/k\_r4: 0.0301642 FCepsilonRI/k\_r6: 0.00398645 Ssq[1.22238094e-201.22238094e-20]

15.FCepsilonRI/k\_f1: 48.9162 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 3.75162 FCepsilonRI/k\_f6: 10.2044 FCepsilonRI/k\_f7: 1.88959 FCepsilonRI/k\_r1: 0.0036682 FCepsilonRI/k\_r4: 40.2246 FCepsilonRI/k\_r6: 0.418861 Ssq[3.15544362e-30 3.15544362e-30]

16.FCepsilonRI/k\_f1: 56.4876 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 3.75162 FCepsilonRI/k\_f6: 10.2044 FCepsilonRI/k\_f7: 1.88959 FCepsilonRI/k\_r1: 0.0036682 FCepsilonRI/k\_r4: 40.2246 FCepsilonRI/k\_r6: 0.418861 Ssq[1.45264713e-24 1.45264713e-24]

17.FCepsilonRI/k\_f1: 48.9162 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 3.75162 FCepsilonRI/k\_f6: 1.41063 FCepsilonRI/k\_f7: 1.88959 FCepsilonRI/k\_r1: 0.0036682 FCepsilonRI/k\_r4: 40.2246 FCepsilonRI/k\_r6: 0.418861 Ssq[2.90191857e-10 2.90191857e-10]

18.FCepsilonRI/k\_f1: 48.9162 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 3.75162 FCepsilonRI/k\_f6: 10.2044 FCepsilonRI/k\_f7: 8.5244 FCepsilonRI/k\_r1: 0.0036682 FCepsilonRI/k\_r4: 40.2246 FCepsilonRI/k\_r6: 0.418861 Ssq[7.88860905e-31 7.88860905e-31]

19.FCepsilonRI/k\_f1: 48.9162 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 3.75162 FCepsilonRI/k\_f6: 10.2044 FCepsilonRI/k\_f7: 1.88959 FCepsilonRI/k\_r1: 0.0032929 FCepsilonRI/k\_r4: 40.2246 FCepsilonRI/k\_r6: 0.418861 Ssq[9.71955521e-27 9.71955521e-27]

20.FCepsilonRI/k\_f1: 48.9162 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 3.75162 FCepsilonRI/k\_f6: 10.2044 FCepsilonRI/k\_f7: 1.88959 FCepsilonRI/k\_r1: 0.0036682 FCepsilonRI/k\_r4: 6.96263 FCepsilonRI/k\_r6: 0.418861 Ssq[1.37409475e-22 1.37409475e-22]

21.FCepsilonRI/k\_f1: 48.9162 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 3.75162 FCepsilonRI/k\_f6: 10.2044 FCepsilonRI/k\_f7: 1.88959 FCepsilonRI/k\_r1: 0.0036682 FCepsilonRI/k\_r4: 40.2246 FCepsilonRI/k\_r6: 0.0168107 Ssq[4.82654163e-23 4.82654163e-23]

22.FCepsilonRI/k\_f1: 56.4876 FCepsilonRI/k\_f4: 90.6817 FCepsilonRI/k\_f5: 0.691582 FCepsilonRI/k\_f6: 1.41063 FCepsilonRI/k\_f7: 8.5244 FCepsilonRI/k\_r1: 0.0032929 FCepsilonRI/k\_r4: 6.96263 FCepsilonRI/k\_r6: 0.0168107 Ssq[3.41867967e-07 3.41867967e-07]

23.FCepsilonRI/k\_f1: 56.4876 FCepsilonRI/k\_f4: 6.18734 FCepsilonRI/k\_f5: 3.75162

FCepsilonRI/k\_f6: 1.41063 FCepsilonRI/k\_f7: 8.5244 FCepsilonRI/k\_r1: 0.0032929 FCepsilonRI/k\_r4: 6.96263 FCepsilonRI/k\_r6: 0.0168107 Ssq[1.7995195e-11 1.7995195e-11]

24.FCepsilonRI/k\_f1: 13.3953 FCepsilonRI/k\_f4: 34.3277 FCepsilonRI/k\_f5: 1.58204 FCepsilonRI/k\_f5: 61.665 FCepsilonRI/k\_f7: 51.5127 FCepsilonRI/k\_r1: 0.0317653 FCepsilonRI/k\_r4: 0.182281 FCepsilonRI/k\_r6: 0.0337536 Ssq[2.83989926e-29 2.83989926e-29]

25.FCepsilonRI/k\_f1: 11.5999 FCepsilonRI/k\_f4: 74.0677 FCepsilonRI/k\_f5: 1.58204 FCepsilonRI/k\_f6: 4.15111 FCepsilonRI/k\_f7: 51.5127 FCepsilonRI/k\_r1: 0.0285153 FCepsilonRI/k\_r4: 0.0074821 FCepsilonRI/k\_r6: 0.00571263 Ssq[3.91577661e-193.91577661e-19]

26.FCepsilonRI/k\_f1: 4.23597 FCepsilonRI/k\_f4: 81.4038 FCepsilonRI/k\_f5: 15.8204 FCepsilonRI/k\_f5: 3.46768 FCepsilonRI/k\_f7: 23.4486 FCepsilonRI/k\_r1: 0.00100451 FCepsilonRI/k\_r4: 0.0102504 FCepsilonRI/k\_r6: 0.600234 Ssq[2.18427381e-24 2.18427381e-24]

27.FCepsilonRI/k\_f1: 3.6682 FCepsilonRI/k\_f4: 81.4038 FCepsilonRI/k\_f5: 15.8204 FCepsilonRI/k\_f5: 3.46768 FCepsilonRI/k\_f7: 23.4486 FCepsilonRI/k\_r1: 0.00100451 FCepsilonRI/k\_r4: 0.0102504 FCepsilonRI/k\_r6: 0.600234 Ssq[3.63566289e-22 3.63566289e-22] 28.FCepsilonRI/k\_f1: 4.23597 FCepsilonRI/k\_f4: 5.55429 FCepsilonRI/k\_f5: 15.8204 FCepsilonRI/k\_f5: 3.46768 FCepsilonRI/k\_f7: 23.4486 FCepsilonRI/k\_r1: 0.00100451 FCepsilonRI/k\_r4: 0.0102504 FCepsilonRI/k\_r6: 0.600234 Ssq[6.76648457e-21 6.76648457e-21]

29.FCepsilonRI/k\_f1: 4.23597 FCepsilonRI/k\_f4: 81.4038 FCepsilonRI/k\_f5: 92.224 FCepsilonRI/k\_f6: 3.46768 FCepsilonRI/k\_f7: 23.4486 FCepsilonRI/k\_r1: 0.00100451 FCepsilonRI/k\_r4: 0.0102504 FCepsilonRI/k\_r6: 0.600234 Ssq[6.91168371e-26 6.91168371e-26]

30.FCepsilonRI/k\_f1: 4.23597 FCepsilonRI/k\_f4: 81.4038 FCepsilonRI/k\_f5: 15.8204 FCepsilonRI/k\_f6: 3.46768 FCepsilonRI/k\_f7: 2.89677 FCepsilonRI/k\_r1: 0.00100451 FCepsilonRI/k\_r4: 0.0102504 FCepsilonRI/k\_r6: 0.600234 Ssq[5.19285516e-19 5.19285516e-19]

31.FCepsilonRI/k\_f1: 4.23597 FCepsilonRI/k\_f4: 81.4038 FCepsilonRI/k\_f5: 15.8204 FCepsilonRI/k\_f5: 3.46768 FCepsilonRI/k\_f7: 23.4486 FCepsilonRI/k\_r1: 0.0901733 FCepsilonRI/k\_r4: 0.0102504 FCepsilonRI/k\_r6: 0.600234 Ssq[3.73898049e-20 3.73898049e-20]

32.FCepsilonRI/k\_f1: 4.23597 FCepsilonRI/k\_f4: 81.4038 FCepsilonRI/k\_f5: 15.8204 FCepsilonRI/k\_f6: 3.46768 FCepsilonRI/k\_f7: 23.4486 FCepsilonRI/k\_r1: 0.00100451 FCepsilonRI/k\_r4: 42.0749 FCepsilonRI/k\_r6: 0.600234 Ssq[3.0589111e-20 3.0589111e-20]

33.FCepsilonRI/k\_f1: 4.23597 FCepsilonRI/k\_f4: 81.4038 FCepsilonRI/k\_f5: 15.8204 FCepsilonRI/k\_f6: 3.46768 FCepsilonRI/k\_f7: 23.4486 FCepsilonRI/k\_r1: 0.00100451 FCepsilonRI/k\_r4: 0.0102504 FCepsilonRI/k\_r6: 0.101586 Ssq[2.3304837e-20 2.3304837e-20]

34.FCepsilonRI/k\_f1: 3.6682 FCepsilonRI/k\_f4: 5.55429 FCepsilonRI/k\_f5: 92.224 FCepsilonRI/k\_f5: 92.224 FCepsilonRI/k\_f6: 3.46768 FCepsilonRI/k\_f7: 2.89677 FCepsilonRI/k\_r1: 0.0901733 FCepsilonRI/k\_r4: 42.0749 FCepsilonRI/k\_r6: 0.101586 Ssq[6.11801147e-20 6.11801147e-20]

35.FCepsilonRI/k\_f1: 7.53274 FCepsilonRI/k\_f4: 9.40036 FCepsilonRI/k\_f5: 5.00286 FCepsilonRI/k\_f6: 0.822317 FCepsilonRI/k\_f7: 98.882 FCepsilonRI/k\_r1: 0.00564876 FCepsilonRI/k\_r4: 0.768674 FCepsilonRI/k\_r6: 0.00800424 Ssq[2.60516451e-09 2.60516451e-09]

36.FCepsilonRI/k\_f1: 65.2309 FCepsilonRI/k\_f4: 9.40036 FCepsilonRI/k\_f5: 5.00286 FCepsilonRI/k\_f6: 0.822317 FCepsilonRI/k\_f7: 98.882 FCepsilonRI/k\_r1: 0.00564876 FCepsilonRI/k\_r4: 0.768674 FCepsilonRI/k\_r6: 0.00800424 Ssq[2.62352062e-09 2.62352062e-09]

37.FCepsilonRI/k\_f1: 7.53274 FCepsilonRI/k\_f4: 48.0982 FCepsilonRI/k\_f5: 5.00286 FCepsilonRI/k\_f6: 0.822317 FCepsilonRI/k\_f7: 98.882 FCepsilonRI/k\_r1: 0.00564876 FCepsilonRI/k\_r4: 0.768674 FCepsilonRI/k\_r6: 0.00800424 Ssq[3.18138982e-10 3.18138982e-10]

38.FCepsilonRI/k\_f1: 7.53274 FCepsilonRI/k\_f4: 9.40036 FCepsilonRI/k\_f5: 5.00286 FCepsilonRI/k\_f6: 0.822317 FCepsilonRI/k\_f7: 98.882 FCepsilonRI/k\_r1: 0.0160353 FCepsilonRI/k\_r4: 0.768674 FCepsilonRI/k\_r6: 0.00800424 Ssq[2.66170979e-09 2.66170979e-09]

39.FCepsilonRI/k\_f1: 7.53274 FCepsilonRI/k\_f4: 9.40036 FCepsilonRI/k\_f5: 5.00286 FCepsilonRI/k\_f6: 0.822317 FCepsilonRI/k\_f7: 98.882 FCepsilonRI/k\_r1: 0.00564876 FCepsilonRI/k\_r4: 9.97754 FCepsilonRI/k\_r6: 0.00800424 Ssq[5.57380022e-08 5.57380022e-08]

40.FCepsilonRI/k\_f1: 7.53274 FCepsilonRI/k\_f4: 9.40036 FCepsilonRI/k\_f5: 5.00286 FCepsilonRI/k\_f6: 0.822317 FCepsilonRI/k\_f7: 98.882 FCepsilonRI/k\_r1: 0.00564876 FCepsilonRI/k\_r4: 0.768674 FCepsilonRI/k\_r6: 0.02409 Ssq[2.72340003e-09 2.72340003e-09]

41.FCepsilonRI/k\_f1: 75.3274 FCepsilonRI/k\_f4: 8.55321 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f6: 17.5051 FCepsilonRI/k\_f7: 0.686932 FCepsilonRI/k\_r1: 0.00160353 FCepsilonRI/k\_r4: 0.0315517 FCepsilonRI/k\_r6: 7.61791 Ssq[5.33566783e-08 5.33566783e-08]

42.FCepsilonRI/k\_f1: 6.52309 FCepsilonRI/k\_f4: 52.8621 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f6: 17.5051 FCepsilonRI/k\_f7: 0.686932 FCepsilonRI/k\_r1: 0.00160353 FCepsilonRI/k\_r4: 0.0315517 FCepsilonRI/k\_r6: 7.61791 Ssq[3.04776291e-09 3.04776291e-09]

43.FCepsilonRI/k\_f1: 6.52309 FCepsilonRI/k\_f4: 8.55321 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f7: 0.686932 FCepsilonRI/k\_f7: 0.0564876 FCepsilonRI/k\_r4: 0.0315517 FCepsilonRI/k\_r6: 7.61791 Ssq[5.35305033e-08 5.35305033e-08]

44.FCepsilonRI/k\_f1: 6.52309 FCepsilonRI/k\_f4: 8.55321 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f6: 17.5051 FCepsilonRI/k\_f7: 0.686932 FCepsilonRI/k\_r1: 0.00160353 FCepsilonRI/k\_r4: 0.00243076 FCepsilonRI/k\_r6: 7.61791 Ssq[5.32152659e-08 5.32152659e-08]

45.FCepsilonRI/k\_f1: 6.52309 FCepsilonRI/k\_f4: 8.55321 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f6: 17.5051 FCepsilonRI/k\_f7: 0.686932 FCepsilonRI/k\_r1: 0.00160353 FCepsilonRI/k\_r4: 0.0315517 FCepsilonRI/k\_r6: 2.53116 Ssq[1.61545619e-21 1.61545619e-21]

46.FCepsilonRI/k\_f1: 6.52309 FCepsilonRI/k\_f4: 8.55321 FCepsilonRI/k\_f5: 29.1638 FCepsilonRI/k\_f6: 17.5051 FCepsilonRI/k\_f7: 0.686932 FCepsilonRI/k\_r1: 0.00160353 FCepsilonRI/k\_r4: 0.0315517 FCepsilonRI/k\_r6: 7.61791 Ssq[5.34167049e-08 5.34167049e-08] 47.FCepsilonRI/k\_f1: 56.4876 FCepsilonRI/k\_f4: 4.9192 FCepsilonRI/k\_f5: 8.89649 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 11.4187 FCepsilonRI/k\_r1: 0.0423597 FCepsilonRI/k\_r4: 6.65644 FCepsilonRI/k\_r6: 92.4316 Ssq[2.70111999e-18 2.70111999e-18]

48.FCepsilonRI/k\_f1: 56.4876 FCepsilonRI/k\_f4: 91.9134 FCepsilonRI/k\_f5: 16.4 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 5.94859 FCepsilonRI/k\_r1: 0.0120248 FCepsilonRI/k\_r4: 4.85874 FCepsilonRI/k\_r6: 0.00278186 Ssq[3.86541844e-29 3.86541844e-29]

49.FCepsilonRI/k\_f1: 2.75076 FCepsilonRI/k\_f4: 4.9192 FCepsilonRI/k\_f5: 16.4 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 5.94859 FCepsilonRI/k\_r1: 0.0120248 FCepsilonRI/k\_r4: 4.85874 FCepsilonRI/k\_r6: 0.00278186 Ssq[7.88860905e-29 7.88860905e-29]

50.FCepsilonRI/k\_f1: 2.75076 FCepsilonRI/k\_f4: 91.9134 FCepsilonRI/k\_f5: 8.89649 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 5.94859 FCepsilonRI/k\_r1: 0.0120248 FCepsilonRI/k\_r4: 4.85874 FCepsilonRI/k\_r6: 0.00278186 Ssq[7.88860905e-31 7.88860905e-31]

51.FCepsilonRI/k\_f1: 2.75076 FCepsilonRI/k\_f4: 91.9134 FCepsilonRI/k\_f5: 16.4 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 11.4187 FCepsilonRI/k\_r1: 0.0120248 FCepsilonRI/k\_r4: 4.85874 FCepsilonRI/k\_r6: 0.00278186

Ssq[1.97215226e-291.97215226e-29]

52.FCepsilonRI/k\_f1: 2.75076 FCepsilonRI/k\_f4: 91.9134 FCepsilonRI/k\_f5: 16.4 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 5.94859 FCepsilonRI/k\_r1: 0.0423597 FCepsilonRI/k\_r4: 4.85874 FCepsilonRI/k\_r6: 0.00278186 Ssq[3.86541844e-29 3.86541844e-29]

53.FCepsilonRI/k\_f1: 2.75076 FCepsilonRI/k\_f4: 91.9134 FCepsilonRI/k\_f5: 16.4 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 5.94859 FCepsilonRI/k\_r1: 0.0120248 FCepsilonRI/k\_r4: 6.65644 FCepsilonRI/k\_r6: 0.00278186 Ssq[9.54521695e-29 9.54521695e-29]

54.FCepsilonRI/k\_f1: 2.75076 FCepsilonRI/k\_f4: 91.9134 FCepsilonRI/k\_f5: 16.4 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 5.94859 FCepsilonRI/k\_r1: 0.0120248 FCepsilonRI/k\_r4: 4.85874 FCepsilonRI/k\_r6: 92.4316 Ssq[3.57584162e-18 3.57584162e-18]

55.FCepsilonRI/k\_f1: 2.75076 FCepsilonRI/k\_f4: 91.9134 FCepsilonRI/k\_f5: 16.4 FCepsilonRI/k\_f6: 35.9471 FCepsilonRI/k\_f7: 5.94859 FCepsilonRI/k\_r1: 0.0120248 FCepsilonRI/k\_r4: 4.85874 FCepsilonRI/k\_r6: 0.00278186 Ssq[7.09974815e-30 7.09974815e-30]

56.FCepsilonRI/k\_f1: 1.54687 FCepsilonRI/k\_f4: 25.1698 FCepsilonRI/k\_f5: 51.8613 FCepsilonRI/k\_f6: 30.0289 FCepsilonRI/k\_f7: 25.085 FCepsilonRI/k\_r1: 0.00676205 FCepsilonRI/k\_r4: 0.00364354 FCepsilonRI/k\_r6: 3.70968 Ssq[1.26217745e-29 1.26217745e-29]

57.FCepsilonRI/k\_f1: 31.7653 FCepsilonRI/k\_f4: 17.9636 FCepsilonRI/k\_f5: 28.1332 FCepsilonRI/k\_f5: 28.1332 FCepsilonRI/k\_f7: 48.1523 FCepsilonRI/k\_r1: 0.0238206 FCepsilonRI/k\_r4: 0.0887651 FCepsilonRI/k\_r6: 0.0693139 Ssq[1.26217745e-29 1.26217745e-29]

58.FCepsilonRI/k\_f1: 1.00677 FCepsilonRI/k\_f4: 32.1425 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 3.40969 FCepsilonRI/k\_r1: 0.0047507 FCepsilonRI/k\_r4: 0.00458798 FCepsilonRI/k\_r6: 19.2605 Ssq[7.88860905e-31 7.88860905e-31]

59.FCepsilonRI/k\_f1: 24.7488 FCepsilonRI/k\_f4: 11.6064 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 3.40969 FCepsilonRI/k\_r1: 0.0047507 FCepsilonRI/k\_r4: 0.00458798 FCepsilonRI/k\_r6: 19.2605 Ssq[1.17928547e-20 1.17928547e-20]

60.FCepsilonRI/k\_f1: 24.7488 FCepsilonRI/k\_f4: 32.1425 FCepsilonRI/k\_f5: 0.618039 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 3.40969 FCepsilonRI/k\_r1: 0.0047507 FCepsilonRI/k\_r4: 0.00458798 FCepsilonRI/k\_r6: 19.2605 Ssq[1.73989226e-18 1.73989226e-18]

61.FCepsilonRI/k\_f1: 24.7488 FCepsilonRI/k\_f4: 32.1425 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 0.610442 FCepsilonRI/k\_r1: 0.0047507 FCepsilonRI/k\_r4: 0.00458798 FCepsilonRI/k\_r6: 19.2605 Ssq[2.55843815e-06 2.55843815e-06]

62.FCepsilonRI/k\_f1: 24.7488 FCepsilonRI/k\_f4: 32.1425 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 3.40969 FCepsilonRI/k\_r1: 0.0488064 FCepsilonRI/k\_r4: 0.00458798 FCepsilonRI/k\_r6: 19.2605 Ssq[5.04870979e-29 5.04870979e-29]

63.FCepsilonRI/k\_f1: 24.7488 FCepsilonRI/k\_f4: 32.1425 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 3.40969 FCepsilonRI/k\_r1: 0.0047507 FCepsilonRI/k\_r4: 0.016162 FCepsilonRI/k\_r6: 19.2605 Ssq[1.19985744e-27 1.19985744e-27]

64.FCepsilonRI/k\_f1: 24.7488 FCepsilonRI/k\_f1: 32.1425 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 3.40969 FCepsilonRI/k\_r1: 0.0047507 FCepsilonRI/k\_r4: 0.00458798 FCepsilonRI/k\_r6: 0.864991 Ssq[4.17307419e-28 4.17307419e-28]

65.FCepsilonRI/k\_f1: 24.7488 FCepsilonRI/k\_f4: 32.1425 FCepsilonRI/k\_f5: 19.3691 FCepsilonRI/k\_f6: 40.4514 FCepsilonRI/k\_f7: 3.40969 FCepsilonRI/k\_r1: 0.0047507 FCepsilonRI/k\_r4: 0.00458798 FCepsilonRI/k\_r6: 19.2605 Ssq[1.29934049e-19 1.29934049e-19] 66.FCepsilonRI/k\_f1: 3.18368 FCepsilonRI/k\_f4: 27.5231 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f6: 4.8315 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.0154339 FCepsilonRI/k\_r4: 90.8858 FCepsilonRI/k\_r6: 15.382 Ssq[1.20442332e-15 1.20442332e-15]

67.FCepsilonRI/k\_f1: 78.2627 FCepsilonRI/k\_f4: 27.5231 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f6: 4.8315 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.0154339 FCepsilonRI/k\_r4: 90.8858 FCepsilonRI/k\_r6: 15.382 Ssq[4.73978457e-164.73978457e-16]

68.FCepsilonRI/k\_f1: 3.18368 FCepsilonRI/k\_f4: 76.222 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f6: 4.8315 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.0154339 FCepsilonRI/k\_r4: 90.8858 FCepsilonRI/k\_r6: 15.382 Ssq[1.5939609e-18 1.5939609e-18]

69.FCepsilonRI/k\_f1: 3.18368 FCepsilonRI/k\_f4: 27.5231 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f6: 2.27475 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.0154339 FCepsilonRI/k\_r4: 90.8858 FCepsilonRI/k\_r6: 15.382 Ssq[2.42546642e-07 2.42546642e-07]

70.FCepsilonRI/k\_f1: 3.18368 FCepsilonRI/k\_f4: 27.5231 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f6: 4.8315 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.015023 FCepsilonRI/k\_r4: 90.8858 FCepsilonRI/k\_r6: 15.382 Ssq[6.27501046e-16 6.27501046e-16]

71.FCepsilonRI/k\_f1: 3.18368 FCepsilonRI/k\_f4: 27.5231 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f6: 4.8315 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.0154339 FCepsilonRI/k\_r4: 0.081587 FCepsilonRI/k\_r6: 15.382 Ssq[6.37312305e-166.37312305e-16]

72.FCepsilonRI/k\_f1: 3.18368 FCepsilonRI/k\_f4: 27.5231 FCepsilonRI/k\_f5: 6.18039 FCepsilonRI/k\_f6: 4.8315 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.0154339 FCepsilonRI/k\_r4: 90.8858 FCepsilonRI/k\_r6: 0.00342506 Ssq[1.44132215e-18 1.44132215e-18]

73.FCepsilonRI/k\_f1: 78.2627 FCepsilonRI/k\_f4: 76.222 FCepsilonRI/k\_f5: 1.93691 FCepsilonRI/k\_f6: 2.27475 FCepsilonRI/k\_f7: 10.8554 FCepsilonRI/k\_r1: 0.015023 FCepsilonRI/k\_r4: 0.081587 FCepsilonRI/k\_r6: 0.00342506 Ssq[1.96094275e-17 1.96094275e-17]

74.FCepsilonRI/k\_f1: 5.66148 FCepsilonRI/k\_f4: 3.17832 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 1.14573 FCepsilonRI/k\_f7: 45.7767 FCepsilonRI/k\_r1: 0.0867914 FCepsilonRI/k\_r4: 0.0681547 FCepsilonRI/k\_r6: 0.205122 Ssq[8.87719195e-12 8.87719195e-12]

75.FCepsilonRI/k\_f1: 4.40103 FCepsilonRI/k\_f4: 3.17832 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 1.14573 FCepsilonRI/k\_f7: 45.7767 FCepsilonRI/k\_r1: 0.0867914 FCepsilonRI/k\_r4: 0.0681547 FCepsilonRI/k\_r6: 0.205122 Ssq[8.97145383e-12 8.97145383e-12]

76.FCepsilonRI/k\_f1: 5.66148 FCepsilonRI/k\_f4: 3.71176 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 1.14573 FCepsilonRI/k\_f7: 45.7767 FCepsilonRI/k\_r1: 0.0867914 FCepsilonRI/k\_r4: 0.0681547 FCepsilonRI/k\_r6: 0.205122 Ssq[3.78804545e-12 3.78804545e-12]

77.FCepsilonRI/k\_f1: 5.66148 FCepsilonRI/k\_f4: 3.17832 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 0.539428 FCepsilonRI/k\_f7: 45.7767 FCepsilonRI/k\_r1: 0.0867914 FCepsilonRI/k\_r4: 0.0681547 FCepsilonRI/k\_r6: 0.205122 Ssq[2.26971278e-05 2.26971278e-05]

78.FCepsilonRI/k\_f1: 5.66148 FCepsilonRI/k\_f4: 3.17832 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 1.14573 FCepsilonRI/k\_f7: 14.3785 FCepsilonRI/k\_r1: 0.0867914 FCepsilonRI/k\_r4: 0.0681547 FCepsilonRI/k\_r6: 0.205122 Ssq[1.79317405e-111.79317405e-11]

79.FCepsilonRI/k\_f1: 5.66148 FCepsilonRI/k\_f4: 3.17832 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 1.14573 FCepsilonRI/k\_f7: 45.7767 FCepsilonRI/k\_r1: 0.0844808 FCepsilonRI/k\_r4: 0.0681547 FCepsilonRI/k\_r6: 0.205122 Ssq[9.67879811e-12 9.67879811e-12]

80.FCepsilonRI/k\_f1: 5.66148 FCepsilonRI/k\_f4: 3.17832

FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 1.14573 FCepsilonRI/k\_f7: 45.7767 FCepsilonRI/k\_r1: 0.0867914 FCepsilonRI/k\_r4: 0.0193473 FCepsilonRI/k\_r6: 0.205122 Ssq[9.67911183e-12 9.67911183e-12]

81.FCepsilonRI/k\_f1: 5.66148 FCepsilonRI/k\_f4: 3.17832 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 1.14573 FCepsilonRI/k\_f7: 45.7767 FCepsilonRI/k\_r1: 0.0867914 FCepsilonRI/k\_r4: 0.0681547 FCepsilonRI/k\_r6: 0.0144434 Ssq[8.53483166e-12 8.53483166e-12]

82.FCepsilonRI/k\_f1: 4.40103 FCepsilonRI/k\_f4: 3.71176 FCepsilonRI/k\_f5: 19.5441 FCepsilonRI/k\_f6: 0.539428 FCepsilonRI/k\_f7: 14.3785 FCepsilonRI/k\_r1: 0.0844808 FCepsilonRI/k\_r4: 0.0193473 FCepsilonRI/k\_r6: 0.0144434 Ssq[1.7878053e-05 1.7878053e-05]

83.FCepsilonRI/k\_f1: 17.9032 FCepsilonRI/k\_f4: 7.53698 FCepsilonRI/k\_f5: 1.95441 FCepsilonRI/k\_f6: 20.3743 FCepsilonRI/k\_f7: 0.808565 FCepsilonRI/k\_r1: 0.00274458 FCepsilonRI/k\_r4: 0.00383262 FCepsilonRI/k\_r6: 3.64764 Ssq[3.42937424e-15 3.42937424e-15]

84.FCepsilonRI/k\_f1: 13.9173 FCepsilonRI/k\_f4: 49.4971 FCepsilonRI/k\_f5: 61.2505 FCepsilonRI/k\_f6: 20.3743 FCepsilonRI/k\_f7: 0.808565 FCepsilonRI/k\_r1: 0.0267152 FCepsilonRI/k\_r4: 0.34405 FCepsilonRI/k\_r6: 0.256843 Ssq[1.61167438e-23 1.61167438e-23] 85.FCepsilonRI/k\_f1: 33.0031 FCepsilonRI/k\_f4: 16.8202 FCepsilonRI/k\_f5: 34.4437 FCepsilonRI/k\_f5: 0.557933 FCepsilonRI/k\_f7: 7.00188 FCepsilonRI/k\_r1: 0.00200335 FCepsilonRI/k\_r4: 0.167541 FCepsilonRI/k\_r6: 2.22417 Ssq[6.84180211e-05 6.84180211e-05]

86.FCepsilonRI/k\_f1: 1.34255 FCepsilonRI/k\_f4: 4.60607 FCepsilonRI/k\_f5: 10.892 FCepsilonRI/k\_f6: 19.6985 FCepsilonRI/k\_f7: 29.5267 FCepsilonRI/k\_r1: 0.0356252 FCepsilonRI/k\_r4: 12.5638 FCepsilonRI/k\_r6: 0.00166789 Ssq[1.54616737e-28 1.54616737e-28]

87.FCepsilonRI/k\_f1: 58.6887 FCepsilonRI/k\_f4: 22.1793 FCepsilonRI/k\_f5: 10.892 FCepsilonRI/k\_f6: 19.6985 FCepsilonRI/k\_f7: 29.5267 FCepsilonRI/k\_r1: 0.0356252 FCepsilonRI/k\_r4: 12.5638 FCepsilonRI/k\_r6: 0.00166789 Ssq[3.15544362e-30 3.15544362e-30]

88.FCepsilonRI/k\_f1: 58.6887 FCepsilonRI/k\_f4: 4.60607 FCepsilonRI/k\_f5: 34.7549 FCepsilonRI/k\_f6: 19.6985 FCepsilonRI/k\_f7: 29.5267 FCepsilonRI/k\_r1: 0.0356252 FCepsilonRI/k\_r4: 12.5638 FCepsilonRI/k\_r6: 0.00166789 Ssq[3.15544362e-30 3.15544362e-30]

89.FCepsilonRI/k\_f1: 58.6887 FCepsilonRI/k\_f4: 4.60607 FCepsilonRI/k\_f5: 10.892 FCepsilonRI/k\_f6: 19.6985 FCepsilonRI/k\_f7: 29.5267 FCepsilonRI/k\_r1: 0.0650843 FCepsilonRI/k\_r4: 12.5638 FCepsilonRI/k\_r6: 0.00166789 Ssq[9.25923594e-24 9.25923594e-24]

90.FCepsilonRI/k\_f1: 58.6887 FCepsilonRI/k\_f4: 4.60607 FCepsilonRI/k\_f5: 10.892 FCepsilonRI/k\_f6: 19.6985 FCepsilonRI/k\_f7: 29.5267 FCepsilonRI/k\_r1: 0.0356252 FCepsilonRI/k\_r4: 44.2584 FCepsilonRI/k\_r6: 0.00166789 Ssq[2.79586722e-22 2.79586722e-22]

91.FCepsilonRI/k\_f1: 58.6887 FCepsilonRI/k\_f4: 4.60607 FCepsilonRI/k\_f5: 10.892 FCepsilonRI/k\_f6: 19.6985 FCepsilonRI/k\_f7: 29.5267 FCepsilonRI/k\_r1: 0.0356252 FCepsilonRI/k\_r4: 12.5638 FCepsilonRI/k\_r6: 7.49052 Ssq[7.88860905e-29 7.88860905e-29]

92.FCepsilonRI/k\_f1: 58.6887 FCepsilonRI/k\_f4: 4.60607 FCepsilonRI/k\_f5: 10.892 FCepsilonRI/k\_f6: 19.6985 FCepsilonRI/k\_f7: 29.5267 FCepsilonRI/k\_r1: 0.0356252 FCepsilonRI/k\_r4: 12.5638 FCepsilonRI/k\_r6: 0.00166789 Ssq[2.27980802e-28 2.27980802e-28]

93.FCepsilonRI/k\_f1: 1.55035 FCepsilonRI/k\_f4: 4.39362 FCepsilonRI/k\_f5: 14.5248 FCepsilonRI/k\_f6: 14.2178 FCepsilonRI/k\_f7: 7.57521 FCepsilonRI/k\_r1: 0.00178228 FCepsilonRI/k\_r4: 3.56653 FCepsilonRI/k\_r6: 0.0339439 Ssq[7.88860905e-29 7.88860905e-29]

94.FCepsilonRI/k\_f1: 4.90264 FCepsilonRI/k\_f4: 58.5898 FCepsilonRI/k\_f5: 14.656 FCepsilonRI/k\_f6: 6.69397 FCepsilonRI/k\_f7: 0.425985 FCepsilonRI/k\_r1: 0.00563607 FCepsilonRI/k\_r4: 0.200561 FCepsilonRI/k\_r6: 0.00190881 Ssq[1.08825569e-11 1.08825569e-11]

95.FCepsilonRI/k\_f1: 4.90264 FCepsilonRI/k\_f4: 44.4329 FCepsilonRI/k\_f5: 1.45248 FCepsilonRI/k\_f5: 6.69397 FCepsilonRI/k\_f7: 20.6046 FCepsilonRI/k\_r1: 0.00308502 FCepsilonRI/k\_r4: 0.0135012 FCepsilonRI/k\_r6: 0.00239011 Ssq[8.27603521e-24 8.27603521e-24]

96.FCepsilonRI/k\_f1: 12.0519 FCepsilonRI/k\_f4: 58.5898 FCepsilonRI/k\_f5: 1.45248 FCepsilonRI/k\_f6: 6.69397 FCepsilonRI/k\_f7: 20.6046 FCepsilonRI/k\_r1: 0.00308502 FCepsilonRI/k\_r4: 0.0135012 FCepsilonRI/k\_r6: 0.00239011 Ssq[1.52723471e-25 1.52723471e-25]

97.FCepsilonRI/k\_f1: 12.0519 FCepsilonRI/k\_f4: 44.4329 FCepsilonRI/k\_f5: 14.656 FCepsilonRI/k\_f5: 6.69397 FCepsilonRI/k\_f7: 20.6046 FCepsilonRI/k\_r1: 0.00308502 FCepsilonRI/k\_r4: 0.0135012 FCepsilonRI/k\_r6: 0.00239011 Ssq[1.19985744e-27 1.19985744e-27]

98.FCepsilonRI/k\_f1: 12.0519 FCepsilonRI/k\_f4: 44.4329 FCepsilonRI/k\_f5: 1.45248 FCepsilonRI/k\_f5: 6.69397 FCepsilonRI/k\_f7: 20.6046 FCepsilonRI/k\_r1: 0.00563607 FCepsilonRI/k\_r4: 0.0135012 FCepsilonRI/k\_r6: 0.00239011 Ssq[2.95521056e-21 2.95521056e-21]

99.FCepsilonRI/k\_f1: 12.0519 FCepsilonRI/k\_f4: 44.4329 FCepsilonRI/k\_f5: 1.45248 FCepsilonRI/k\_f6: 6.69397 FCepsilonRI/k\_f7: 20.6046 FCepsilonRI/k\_r1: 0.00308502 FCepsilonRI/k\_r4: 0.200561 FCepsilonRI/k\_r6: 0.00239011 Ssq[2.25573277e-24 2.25573277e-24]

100.FCepsilonRI/k\_f1: 12.0519 FCepsilonRI/k\_f4: 44.4329 FCepsilonRI/k\_f5: 1.45248 FCepsilonRI/k\_f5: 6.69397 FCepsilonRI/k\_f7: 20.6046 FCepsilonRI/k\_r1: 0.00308502 FCepsilonRI/k\_r4: 0.0135012 FCepsilonRI/k\_r6: 0.00190881 Ssq[3.15544362e-30 3.15544362e-30]

101.FCepsilonRI/k\_f1: 12.0519 FCepsilonRI/k\_f4: 44.4329 FCepsilonRI/k\_f5: 1.45248 FCepsilonRI/k\_f6: 6.69397 FCepsilonRI/k\_f7: 20.6046 FCepsilonRI/k\_r1: 0.00308502 FCepsilonRI/k\_r4: 0.0135012 FCepsilonRI/k\_r6: 0.00239011 Ssq[1.31046722e-21 1.31046722e-21]

102.FCepsilonRI/k\_f1: 8.71826 FCepsilonRI/k\_f4: 16.0444 FCepsilonRI/k\_f5: 4.63464 FCepsilonRI/k\_f6: 28.2282 FCepsilonRI/k\_f7: 1.79636 FCepsilonRI/k\_f7: 0.00100225 FCepsilonRI/k\_r4: 15.0399 FCepsilonRI/k\_r6: 2.54543 Ssq[7.37655155e-25 7.37655155e-25]

103.FCepsilonRI/k\_f1: 21.4316 FCepsilonRI/k\_f4: 28.8539 FCepsilonRI/k\_f5: 4.59314 FCepsilonRI/k\_f6: 3.37157 FCepsilonRI/k\_f7: 86.889 FCepsilonRI/k\_r1: 0.0548603 FCepsilonRI/k\_r4: 1.01244 FCepsilonRI/k\_r6: 3.18726 Ssq[2.09287049e-16 2.09287049e-16] 104.FCepsilonRI/k\_f1: 2.75696 FCepsilonRI/k\_f4: 38.0472 FCepsilonRI/k\_f5: 0.463464 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 31.9444 FCepsilonRI/k\_r1: 0.0031694 FCepsilonRI/k\_r4: 0.00267452 FCepsilonRI/k\_r6: 45.2649 Ssq[3.69756856e-22 3.69756856e-22]

105.FCepsilonRI/k\_f1: 6.77727 FCepsilonRI/k\_f4: 38.0472 FCepsilonRI/k\_f5: 0.463464 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 31.9444 FCepsilonRI/k\_r1: 0.0031694 FCepsilonRI/k\_r4: 0.00267452 FCepsilonRI/k\_r6: 45.2649 Ssq[1.33317493e-28 1.33317493e-28]

106.FCepsilonRI/k\_f1: 2.75696 FCepsilonRI/k\_f4: 12.1676 FCepsilonRI/k\_f5: 0.463464 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 31.9444 FCepsilonRI/k\_r1: 0.0031694 FCepsilonRI/k\_r4: 0.00267452 FCepsilonRI/k\_r6: 45.2649 Ssq[1.4101433e-24 1.4101433e-24]

107.FCepsilonRI/k\_f1: 2.75696 FCepsilonRI/k\_f4: 38.0472 FCepsilonRI/k\_f5: 0.459314 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 31.9444 FCepsilonRI/k\_r1: 0.0031694 FCepsilonRI/k\_r4: 0.00267452 FCepsilonRI/k\_r6: 45.2649 Ssq[2.56300908e-27 2.56300908e-27]

108.FCepsilonRI/k\_f1: 2.75696 FCepsilonRI/k\_f4: 38.0472 FCepsilonRI/k\_f5: 0.463464 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 4.88613 FCepsilonRI/k\_r1: 0.0031694 FCepsilonRI/k\_r4: 0.00267452 FCepsilonRI/k\_r6: 45.2649

Ssq[1.86000225e-11 1.86000225e-11]

109.FCepsilonRI/k\_f1: 2.75696 FCepsilonRI/k\_f4: 38.0472 FCepsilonRI/k\_f5: 0.463464 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 31.9444 FCepsilonRI/k\_r1: 0.0173483 FCepsilonRI/k\_r4: 0.00267452 FCepsilonRI/k\_r6: 45.2649 Ssq[2.55590933e-28 2.55590933e-28]

110.FCepsilonRI/k\_f1: 2.75696 FCepsilonRI/k\_f4: 38.0472 FCepsilonRI/k\_f5: 0.463464 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 31.9444 FCepsilonRI/k\_r1: 0.0031694 FCepsilonRI/k\_r4: 0.0569339 FCepsilonRI/k\_r6: 45.2649 Ssq[1.33317493e-28 1.33317493e-28]

111.FCepsilonRI/k\_f1: 2.75696 FCepsilonRI/k\_f4: 38.0472 FCepsilonRI/k\_f5: 0.463464 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 31.9444 FCepsilonRI/k\_r1: 0.0031694 FCepsilonRI/k\_r4: 0.00267452 FCepsilonRI/k\_r6: 56.6785 Ssq[3.12396807e-26 3.12396807e-26]

112.FCepsilonRI/k\_f1: 6.77727 FCepsilonRI/k\_f4: 12.1676 FCepsilonRI/k\_f5: 0.459314 FCepsilonRI/k\_f6: 59.9559 FCepsilonRI/k\_f7: 4.88613 FCepsilonRI/k\_r1: 0.0173483 FCepsilonRI/k\_r4: 0.0569339 FCepsilonRI/k\_r6: 56.6785 Ssq[3.7629008e-06 3.7629008e-06]

113.FCepsilonRI/k\_f1: 2.06742 FCepsilonRI/k\_f4: 8.396 FCepsilonRI/k\_f5: 82.4168 FCepsilonRI/k\_f6: 1.64184 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00751582 FCepsilonRI/k\_r4: 7.32396 FCepsilonRI/k\_r6: 5.22711 Ssq[8.47840165e-15 8.47840165e-15]

114.FCepsilonRI/k\_f1: 2.06742 FCepsilonRI/k\_f4: 55.1385 FCepsilonRI/k\_f5: 0.816788 FCepsilonRI/k\_f6: 0.773007 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00130094 FCepsilonRI/k\_r4: 2.07908 FCepsilonRI/k\_r6: 0.00490815 Ssq[2.8092583e-05 2.8092583e-05]

115.FCepsilonRI/k\_f1: 2.85795 FCepsilonRI/k\_f4: 8.396 FCepsilonRI/k\_f5: 0.816788 FCepsilonRI/k\_f6: 0.773007 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00130094 FCepsilonRI/k\_r4: 2.07908 FCepsilonRI/k\_r6: 0.00490815 Ssq[0.00354825 0.00354825]

116.FCepsilonRI/k\_f1: 2.85795 FCepsilonRI/k\_f4: 55.1385 FCepsilonRI/k\_f5: 82.4168 FCepsilonRI/k\_f5: 82.4168 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00130094 FCepsilonRI/k\_r4: 2.07908 FCepsilonRI/k\_r6: 0.00490815 Ssq[4.71501571e-10 4.71501571e-10]

117.FCepsilonRI/k\_f1: 2.85795 FCepsilonRI/k\_f4: 55.1385 FCepsilonRI/k\_f5: 0.816788 FCepsilonRI/k\_f5: 1.64184 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00130094 FCepsilonRI/k\_r4: 2.07908 FCepsilonRI/k\_r6: 0.00490815 Ssq[3.97293324e-12 3.97293324e-12]

118.FCepsilonRI/k\_f1: 2.85795 FCepsilonRI/k\_f4: 55.1385 FCepsilonRI/k\_f5: 0.816788 FCepsilonRI/k\_f6: 0.773007 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00751582 FCepsilonRI/k\_r4: 2.07908 FCepsilonRI/k\_r6: 0.00490815 Ssq[2.80565546e-05 2.80565546e-05]

119.FCepsilonRI/k\_f1: 2.85795 FCepsilonRI/k\_f4: 55.1385 FCepsilonRI/k\_f5: 0.816788 FCepsilonRI/k\_f6: 0.773007 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00130094 FCepsilonRI/k\_r4: 7.32396 FCepsilonRI/k\_r6: 0.00490815 Ssq[0.0001586 0.0001586]

120.FCepsilonRI/k\_f1: 2.85795 FCepsilonRI/k\_f4: 55.1385 FCepsilonRI/k\_f5: 0.816788 FCepsilonRI/k\_f6: 0.773007 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00130094 FCepsilonRI/k\_r4: 2.07908 FCepsilonRI/k\_r6: 5.22711 Ssq[0.00013054 0.00013054]

121.FCepsilonRI/k\_f1: 2.85795 FCepsilonRI/k\_f4: 55.1385 FCepsilonRI/k\_f5: 0.816788 FCepsilonRI/k\_f5: 0.773007 FCepsilonRI/k\_f7: 42.3121 FCepsilonRI/k\_r1: 0.00130094 FCepsilonRI/k\_r4: 2.07908 FCepsilonRI/k\_r6: 0.00490815 Ssq[2.80551691e-05 2.80551691e-05]

122.FCepsilonRI/k\_f1: 65.3777 FCepsilonRI/k\_f4: 3.54056 FCepsilonRI/k\_f5: 8.24168 FCepsilonRI/k\_f6: 29.1966 FCepsilonRI/k\_f7: 0.874771 FCepsilonRI/k\_r1: 0.0237671 FCepsilonRI/k\_r4: 0.411857 FCepsilonRI/k\_r6: 0.293942 Ssq[2.99964359e-24 2.99964359e-24]

123.FCepsilonRI/k\_f1: 90.3763 FCepsilonRI/k\_f4: 3.54056 FCepsilonRI/k\_f5: 8.24168 FCepsilonRI/k\_f6: 29.1966 FCepsilonRI/k\_f7: 0.874771 FCepsilonRI/k\_r1: 0.0237671 FCepsilonRI/k\_r4: 0.411857 FCepsilonRI/k\_r6: 0.293942 Ssq[1.47220378e-25 1.47220378e-25]

124.FCepsilonRI/k\_f1: 65.3777 FCepsilonRI/k\_f4: 4.13481 FCepsilonRI/k\_f5: 8.24168 FCepsilonRI/k\_f6: 29.1966 FCepsilonRI/k\_f7: 0.874771 FCepsilonRI/k\_r1: 0.0237671 FCepsilonRI/k\_r4: 0.411857 FCepsilonRI/k\_r6: 0.293942 Ssq[1.04036756e-22 1.04036756e-22]

125.FCepsilonRI/k\_f1: 65.3777 FCepsilonRI/k\_f4: 3.54056 FCepsilonRI/k\_f5: 8.16788 FCepsilonRI/k\_f5: 29.1966 FCepsilonRI/k\_f7: 0.874771 FCepsilonRI/k\_r1: 0.0237671 FCepsilonRI/k\_r4: 0.411857 FCepsilonRI/k\_r6: 0.293942 Ssq[1.26217745e-29 1.26217745e-29]

126.FCepsilonRI/k\_f1: 65.3777 FCepsilonRI/k\_f4: 3.54056 FCepsilonRI/k\_f5: 8.24168 FCepsilonRI/k\_f5: 29.1966 FCepsilonRI/k\_f7: 2.37939 FCepsilonRI/k\_r1: 0.0237671 FCepsilonRI/k\_r4: 0.411857 FCepsilonRI/k\_r6: 0.293942 Ssq[2.27980802e-28 2.27980802e-28]

127.FCepsilonRI/k\_f1: 65.3777 FCepsilonRI/k\_f4: 3.54056 FCepsilonRI/k\_f5: 8.24168 FCepsilonRI/k\_f5: 29.1966 FCepsilonRI/k\_f7: 0.874771 FCepsilonRI/k\_r1: 0.00411394 FCepsilonRI/k\_r4: 0.411857 FCepsilonRI/k\_r6: 0.293942 Ssq[7.62994156e-26 7.62994156e-26] 128.FCepsilonRI/k\_f1: 65.3777 FCepsilonRI/k\_f4: 3.54056 FCepsilonRI/k\_f5: 8.24168 FCepsilonRI/k\_f5: 0.874771 FCepsilonRI/k\_f7: 0.874771 FCepsilonRI/k\_r1: 0.0237671 FCepsilonRI/k\_r4: 0.116915 FCepsilonRI/k\_r6: 0.293942 Ssq[2.01948392e-28 2.01948392e-28]

129.FCepsilonRI/k\_f1: 65.3777 FCepsilonRI/k\_f4: 3.54056 FCepsilonRI/k\_f5: 8.24168 FCepsilonRI/k\_f6: 29.1966 FCepsilonRI/k\_f7: 0.874771 FCepsilonRI/k\_r1: 0.0237671 FCepsilonRI/k\_r4: 0.411857 FCepsilonRI/k\_r6: 0.0872806 Ssq[3.12396807e-26 3.12396807e-26]

130.FCepsilonRI/k\_f1: 90.3763 FCepsilonRI/k\_f4: 4.13481 FCepsilonRI/k\_f5: 8.16788 FCepsilonRI/k\_f6: 29.1966 FCepsilonRI/k\_f7: 2.37939 FCepsilonRI/k\_r1: 0.00411394 FCepsilonRI/k\_r4: 0.116915 FCepsilonRI/k\_r6: 0.0872806 Ssq[7.88860905e-31 7.88860905e-31]

131.FCepsilonRI/k\_f1: 3.67646 FCepsilonRI/k\_f4: 15.0992 FCepsilonRI/k\_f5: 2.58291 FCepsilonRI/k\_f6: 57.9674 FCepsilonRI/k\_f7: 0.564241 FCepsilonRI/k\_r1: 0.0231344 FCepsilonRI/k\_r4: 8.76741 FCepsilonRI/k\_r6: 6.54513 Ssq[0.00010716 0.00010716]

132.FCepsilonRI/k\_f1: 50.8223 FCepsilonRI/k\_f4: 15.0992 FCepsilonRI/k\_f5: 2.58291 FCepsilonRI/k\_f6: 57.9674 FCepsilonRI/k\_f7: 0.564241 FCepsilonRI/k\_r1: 0.00422646 FCepsilonRI/k\_r4: 8.76741 FCepsilonRI/k\_r6: 6.54513 Ssq[0.00010604 0.00010604] 133.FCepsilonRI/k\_f1: 50.8223 FCepsilonRI/k\_f4: 15.0992 FCepsilonRI/k\_f5: 2.58291 FCepsilonRI/k\_f6: 57.9674 FCepsilonRI/k\_f7: 0.564241 FCepsilonRI/k\_r1: 0.0231344 FCepsilonRI/k\_r4: 1.73679 FCepsilonRI/k\_r6: 6.54513 Ssq[1.23133272e-10 1.23133272e-10]

134.FCepsilonRI/k\_f1: 50.8223 FCepsilonRI/k\_f4: 15.0992 FCepsilonRI/k\_f5: 2.58291 FCepsilonRI/k\_f6: 57.9674 FCepsilonRI/k\_f7: 0.564241 FCepsilonRI/k\_r1: 0.0231344 FCepsilonRI/k\_r4: 8.76741 FCepsilonRI/k\_r6: 0.0697046 Ssq[3.69203708e-20 3.69203708e-20]

135.FCepsilonRI/k\_f1: 50.8223 FCepsilonRI/k\_f4: 15.0992 FCepsilonRI/k\_f5: 2.58291 FCepsilonRI/k\_f6: 57.9674 FCepsilonRI/k\_f7: 0.564241 FCepsilonRI/k\_r1: 0.0231344 FCepsilonRI/k\_r4: 8.76741 FCepsilonRI/k\_r6: 6.54513 Ssq[0.00010605 0.00010605]

136.FCepsilonRI/k\_f1: 36.7646 FCepsilonRI/k\_f4: 30.66 FCepsilonRI/k\_f5: 26.0625 FCepsilonRI/k\_f6: 6.9236 FCepsilonRI/k\_f7: 3.68888 FCepsilonRI/k\_r1: 0.0422646 FCepsilonRI/k\_r4: 0.0054922 FCepsilonRI/k\_r6: 22.0425 Ssq[6.837851e-12 6.837851e-12]

137.FCepsilonRI/k\_f1: 5.08223 FCepsilonRI/k\_f4: 30.66 FCepsilonRI/k\_f5: 26.0625 FCepsilonRI/k\_f6: 6.9236 FCepsilonRI/k\_f7: 3.68888 FCepsilonRI/k\_r1: 0.0422646 FCepsilonRI/k\_r4: 0.0054922 FCepsilonRI/k\_r6: 22.0425 Ssq[7.31491555e-127.31491555e-12] 138.FCepsilonRI/k\_f1: 36.7646 FCepsilonRI/k\_f4: 84.9093 FCepsilonRI/k\_f5: 26.0625 FCepsilonRI/k\_f6: 6.9236 FCepsilonRI/k\_f7: 3.68888 FCepsilonRI/k\_r1: 0.0422646 FCepsilonRI/k\_r4: 0.0054922 FCepsilonRI/k\_r6: 22.0425 Ssq[2.09378037e-12 2.09378037e-12]

139.FCepsilonRI/k\_f1: 36.7646 FCepsilonRI/k\_f4: 30.66 FCepsilonRI/k\_f5: 26.0625 FCepsilonRI/k\_f6: 6.9236 FCepsilonRI/k\_f7: 3.68888 FCepsilonRI/k\_r1: 0.00231344 FCepsilonRI/k\_r4: 0.0054922 FCepsilonRI/k\_r6: 22.0425 Ssq[6.03567335e-12 6.03567335e-12]

140.FCepsilonRI/k\_f1: 36.7646 FCepsilonRI/k\_f4: 30.66 FCepsilonRI/k\_f5: 26.0625 FCepsilonRI/k\_f5: 6.9236 FCepsilonRI/k\_f7: 3.68888 FCepsilonRI/k\_r1: 0.0422646 FCepsilonRI/k\_r4: 0.027725 FCepsilonRI/k\_r6: 22.0425 Ssq[6.23161005e-12 6.23161005e-12]

141.FCepsilonRI/k\_f1: 36.7646 FCepsilonRI/k\_f4: 30.66 FCepsilonRI/k\_f5: 26.0625 FCepsilonRI/k\_f5: 6.9236 FCepsilonRI/k\_f7: 3.68888 FCepsilonRI/k\_r1: 0.0422646 FCepsilonRI/k\_r4: 0.0054922 FCepsilonRI/k\_r6: 0.0206975 Ssq[4.17307419e-28 4.17307419e-28]

142.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 5.16577 FCepsilonRI/k\_f5: 12.6916 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 2.1504 FCepsilonRI/k\_r1: 0.00255403 FCepsilonRI/k\_r4: 52.9811 FCepsilonRI/k\_r6: 0.0138082 Ssq[1.33449777e-23 1.33449777e-23]

143.FCepsilonRI/k\_f1: 4.09548 FCepsilonRI/k\_f4: 5.16577 FCepsilonRI/k\_f5: 12.6916 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 2.1504 FCepsilonRI/k\_r1: 0.00255403 FCepsilonRI/k\_r4: 52.9811 FCepsilonRI/k\_r6: 0.0138082 Ssq[1.26217745e-29 1.26217745e-29]

144.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 30.4539 FCepsilonRI/k\_f5: 12.6916 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 2.1504 FCepsilonRI/k\_r1: 0.00255403 FCepsilonRI/k\_r4: 52.9811 FCepsilonRI/k\_r6: 0.0138082 Ssq[2.84778787e-28 2.84778787e-28]

145.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 5.16577 FCepsilonRI/k\_f5: 7.07309 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 2.1504 FCepsilonRI/k\_r1: 0.00255403 FCepsilonRI/k\_r4: 52.9811 FCepsilonRI/k\_r6: 0.0138082 Ssq[0.00325315 0.00325315]

146.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 5.16577 FCepsilonRI/k\_f5: 12.6916 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 5.84911 FCepsilonRI/k\_r1: 0.00255403 FCepsilonRI/k\_r4: 52.9811 FCepsilonRI/k\_r6: 0.0138082 Ssq[1.81753553e-27 1.81753553e-27]

147.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 5.16577 FCepsilonRI/k\_f5: 12.6916 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 2.1504 FCepsilonRI/k\_r1: 0.0215282 FCepsilonRI/k\_r4: 52.9811 FCepsilonRI/k\_r6: 0.0138082 Ssq[5.1757164e-27 5.1757164e-27]

148.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 5.16577 FCepsilonRI/k\_f5: 12.6916 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 2.1504 FCepsilonRI/k\_r1: 0.00255403 FCepsilonRI/k\_r4: 21.5524 FCepsilonRI/k\_r6: 0.0138082 Ssq[3.64650953e-26 3.64650953e-26]

149.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 5.16577 FCepsilonRI/k\_f5: 12.6916 FCepsilonRI/k\_f6: 50.0849 FCepsilonRI/k\_f7: 2.1504 FCepsilonRI/k\_r1: 0.00255403 FCepsilonRI/k\_r4: 52.9811 FCepsilonRI/k\_r6: 7.83507 Ssq[5.79432853e-22 5.79432853e-22]

150.FCepsilonRI/k\_f1: 4.09548 FCepsilonRI/k\_f4: 30.4539 FCepsilonRI/k\_f5: 7.07309 FCepsilonRI/k\_f5: 50.0849 FCepsilonRI/k\_f7: 5.84911 FCepsilonRI/k\_r1: 0.0215282 FCepsilonRI/k\_r4: 21.5524 FCepsilonRI/k\_r6: 7.83507 Ssq[1.77493704e-28 1.77493704e-28]

151.FCepsilonRI/k\_f1: 22.2167 FCepsilonRI/k\_f4: 46.8967 FCepsilonRI/k\_f5: 2.23671 FCepsilonRI/k\_f6: 5.59189 FCepsilonRI/k\_f7: 24.6655 FCepsilonRI/k\_r1: 0.00382832 FCepsilonRI/k\_r4: 0.287406 FCepsilonRI/k\_r6: 0.104482 Ssq[1.30802478e-23 1.30802478e-23]

152.FCepsilonRI/k\_f1: 7.28291 FCepsilonRI/k\_f4: 3.35455 FCepsilonRI/k\_f5: 2.23671 FCepsilonRI/k\_f6: 5.59189 FCepsilonRI/k\_f7: 24.6655 FCepsilonRI/k\_r1: 0.00382832 FCepsilonRI/k\_r4: 0.287406 FCepsilonRI/k\_r6: 0.104482 Ssq[6.20810505e-22 6.20810505e-22]

153.FCepsilonRI/k\_f1: 7.28291 FCepsilonRI/k\_f4: 46.8967 FCepsilonRI/k\_f5: 2.23671 FCepsilonRI/k\_f6: 5.59189 FCepsilonRI/k\_f7: 24.6655 FCepsilonRI/k\_r1: 0.0454178 FCepsilonRI/k\_r4: 0.287406 FCepsilonRI/k\_r6: 0.104482 Ssq[1.06149123e-26 1.06149123e-26]

154.FCepsilonRI/k\_f1: 7.28291 FCepsilonRI/k\_f4: 46.8967 FCepsilonRI/k\_f5: 2.23671 FCepsilonRI/k\_f6: 5.59189 FCepsilonRI/k\_f7: 24.6655 FCepsilonRI/k\_r1: 0.00382832 FCepsilonRI/k\_r4: 12.5638 FCepsilonRI/k\_r6: 0.104482 Ssq[4.65814476e-24 4.65814476e-24]

155.FCepsilonRI/k\_f1: 7.28291 FCepsilonRI/k\_f4: 46.8967 FCepsilonRI/k\_f5: 2.23671 FCepsilonRI/k\_f6: 5.59189 FCepsilonRI/k\_f7: 24.6655 FCepsilonRI/k\_r1: 0.00382832 FCepsilonRI/k\_r4: 0.287406 FCepsilonRI/k\_r6: 0.00327444 Ssq[7.27595522e-247.27595522e-24]

156.FCepsilonRI/k\_f1: 7.28291 FCepsilonRI/k\_f4: 46.8967 FCepsilonRI/k\_f5: 2.23671 FCepsilonRI/k\_f6: 5.59189 FCepsilonRI/k\_f7: 24.6655 FCepsilonRI/k\_r1: 0.00382832 FCepsilonRI/k\_r4: 0.287406 FCepsilonRI/k\_r6: 0.104482 Ssq[7.94119439e-22 7.94119439e-22] 157.FCepsilonRI/k\_f1: 2.22167 FCepsilonRI/k\_f4: 18.864 FCepsilonRI/k\_f5: 4.01343 FCepsilonRI/k\_f5: 11.877 FCepsilonRI/k\_f7: 0.509941 FCepsilonRI/k\_r1: 0.00454178 FCepsilonRI/k\_r4: 0.0397302 FCepsilonRI/k\_r6: 1.03547 Ssq[1.07713044e-09 1.07713044e-09]

158.FCepsilonRI/k\_f1: 72.8291 FCepsilonRI/k\_f4: 18.864 FCepsilonRI/k\_f5: 4.01343 FCepsilonRI/k\_f5: 11.877 FCepsilonRI/k\_f7: 0.509941 FCepsilonRI/k\_r1: 0.00454178 FCepsilonRI/k\_r4: 0.0397302 FCepsilonRI/k\_r6: 1.03547 Ssq[1.07498391e-09 1.07498391e-09]

159.FCepsilonRI/k\_f1: 2.22167 FCepsilonRI/k\_f4: 8.33955 FCepsilonRI/k\_f5: 4.01343 FCepsilonRI/k\_f5: 11.877 FCepsilonRI/k\_f7: 0.509941 FCepsilonRI/k\_r1: 0.00454178 FCepsilonRI/k\_r4: 0.0397302 FCepsilonRI/k\_r6: 1.03547 Ssq[1.21324993e-07 1.21324993e-07]

160.FCepsilonRI/k\_f1: 2.22167 FCepsilonRI/k\_f4: 18.864 FCepsilonRI/k\_f5: 4.01343 FCepsilonRI/k\_f6: 11.877 FCepsilonRI/k\_f7: 0.509941 FCepsilonRI/k\_f1: 0.0382832 FCepsilonRI/k\_r4: 0.0397302 FCepsilonRI/k\_r6: 1.03547 Ssq[1.08099162e-09 1.08099162e-09]

161.FCepsilonRI/k\_f1: 29.6265 FCepsilonRI/k\_f4: 85.484 FCepsilonRI/k\_f5: 7.137 FCepsilonRI/k\_f6: 0.645741 FCepsilonRI/k\_f7: 4.4159 FCepsilonRI/k\_r1: 0.0605657 FCepsilonRI/k\_r4: 0.00108798 FCepsilonRI/k\_r6: 0.0283555 Ssq[1.32029181e-071.32029181e-07]

162.FCepsilonRI/k\_f1: 3.07118 FCepsilonRI/k\_f4: 15.9365 FCepsilonRI/k\_f5: 1.25779 FCepsilonRI/k\_f6: 0.645741 FCepsilonRI/k\_f7: 2.84833 FCepsilonRI/k\_r1: 0.0510514 FCepsilonRI/k\_r4: 0.590196 FCepsilonRI/k\_r6: 0.0283555 Ssq[0.00020196 0.00020196]

163.FCepsilonRI/k\_f1: 93.6871 FCepsilonRI/k\_f4: 6.41039 FCepsilonRI/k\_f5: 12.5779 FCepsilonRI/k\_f6: 5.78372 FCepsilonRI/k\_f7: 78.5271 FCepsilonRI/k\_r1: 0.0191525 FCepsilonRI/k\_r4: 6.11816 FCepsilonRI/k\_r6: 0.00159454 Ssq[1.28118111e-25 1.28118111e-25]

164.FCepsilonRI/k\_f1: 9.71191 FCepsilonRI/k\_f4: 6.72036 FCepsilonRI/k\_f5: 12.5779 FCepsilonRI/k\_f6: 11.4831 FCepsilonRI/k\_f7: 78.5271 FCepsilonRI/k\_r1: 0.00161439 FCepsilonRI/k\_r4: 10.4953 FCepsilonRI/k\_r6: 0.214557 Ssq[3.15544362e-30 3.15544362e-30]

165.FCepsilonRI/k\_f1: 52.6841 FCepsilonRI/k\_f4: 23.4091 FCepsilonRI/k\_f5: 0.397749 FCepsilonRI/k\_f6: 24.3897 FCepsilonRI/k\_f7: 18.6217 FCepsilonRI/k\_r1: 0.0340586 FCepsilonRI/k\_r4: 0.081587 FCepsilonRI/k\_r6: 2.12636 Ssq[6.45732348e-18 6.45732348e-18]

166.FCepsilonRI/k\_f1: 54.6141 FCepsilonRI/k\_f4: 58.1959 FCepsilonRI/k\_f5: 0.397749 FCepsilonRI/k\_f6: 48.4238 FCepsilonRI/k\_f7: 12.0113

FCepsilonRI/k\_r1: 0.00907837 FCepsilonRI/k\_r4: 0.081587 FCepsilonRI/k\_r6: 0.0508795 Ssq[7.02982352e-257.02982352e-25]

167.FCepsilonRI/k\_f1: 1.72705 FCepsilonRI/k\_f4: 4.36408 FCepsilonRI/k\_f5: 3.97749 FCepsilonRI/k\_f6: 1.37154 FCepsilonRI/k\_f7: 0.675445 FCepsilonRI/k\_r1: 0.00287083 FCepsilonRI/k\_r4: 0.00787039 FCepsilonRI/k\_r6: 0.00286117 Ssq[0.00013549 0.00013549]

168.FCepsilonRI/k\_f1: 4.56226 FCepsilonRI/k\_f4: 26.0771 FCepsilonRI/k\_f5: 53.0407 FCepsilonRI/k\_f6: 0.957099 FCepsilonRI/k\_f7: 1.50062 FCepsilonRI/k\_r1: 0.0124373 FCepsilonRI/k\_r4: 4.26944 FCepsilonRI/k\_r6: 1.48384 Ssq[0.00015616 0.00015616]

169.FCepsilonRI/k\_f1: 4.56226 FCepsilonRI/k\_f4: 26.0771 FCepsilonRI/k\_f5: 0.951734 FCepsilonRI/k\_f6: 0.957099 FCepsilonRI/k\_f7: 1.50062 FCepsilonRI/k\_r1: 0.0124373 FCepsilonRI/k\_r4: 4.26944 FCepsilonRI/k\_r6: 0.00199661 Ssq[0.00295047 0.00295047]

170.FCepsilonRI/k\_f1: 8.41017 FCepsilonRI/k\_f4: 52.2417 FCepsilonRI/k\_f5: 53.0407 FCepsilonRI/k\_f6: 0.957099 FCepsilonRI/k\_f7: 1.98765 FCepsilonRI/k\_f7: 1.00331519 FCepsilonRI/k\_r4: 7.32396 FCepsilonRI/k\_r6: 0.00199661 Ssq[8.99195234e-11 8.99195234e-11]

171.FCepsilonRI/k\_f1: 19.2389 FCepsilonRI/k\_f4: 21.0141 FCepsilonRI/k\_f5: 16.9245 FCepsilonRI/k\_f6: 1.96543 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00165854 FCepsilonRI/k\_r4: 0.493027 FCepsilonRI/k\_r6: 0.722581 Ssq[2.88159643e-18 2.88159643e-18]

172.FCepsilonRI/k\_f1: 3.54654 FCepsilonRI/k\_f4: 21.0141 FCepsilonRI/k\_f5: 16.9245 FCepsilonRI/k\_f6: 1.96543 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00165854 FCepsilonRI/k\_r4: 0.493027 FCepsilonRI/k\_r6: 0.722581 Ssq[7.42143246e-18 7.42143246e-18]

173.FCepsilonRI/k\_f1: 19.2389 FCepsilonRI/k\_f4: 11.5284 FCepsilonRI/k\_f5: 16.9245 FCepsilonRI/k\_f6: 1.96543 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00165854 FCepsilonRI/k\_r4: 0.493027 FCepsilonRI/k\_r6: 0.722581 Ssq[4.75858649e-19 4.75858649e-19]

174.FCepsilonRI/k\_f1: 19.2389 FCepsilonRI/k\_f4: 21.0141 FCepsilonRI/k\_f5: 2.9827 FCepsilonRI/k\_f6: 1.96543 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00165854 FCepsilonRI/k\_r4: 0.493027 FCepsilonRI/k\_r6: 0.722581 Ssq[8.89186667e-17 8.89186667e-17]

175.FCepsilonRI/k\_f1: 19.2389 FCepsilonRI/k\_f4: 21.0141 FCepsilonRI/k\_f5: 16.9245 FCepsilonRI/k\_f6: 33.7916 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00165854 FCepsilonRI/k\_r4: 0.493027 FCepsilonRI/k\_r6: 0.722581 Ssq[9.54521695e-29 9.54521695e-29]

176.FCepsilonRI/k\_f1: 19.2389 FCepsilonRI/k\_f4: 21.0141 FCepsilonRI/k\_f5: 16.9245 FCepsilonRI/k\_f6: 1.96543 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00786154 FCepsilonRI/k\_r4: 0.493027 FCepsilonRI/k\_r6: 0.722581 Ssq[2.89805296e-16 2.89805296e-16]

177.FCepsilonRI/k\_f1: 19.2389 FCepsilonRI/k\_f4: 21.0141 FCepsilonRI/k\_f5: 16.9245 FCepsilonRI/k\_f6: 1.96543 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00165854 FCepsilonRI/k\_r4: 15.0399 FCepsilonRI/k\_r6: 0.722581 Ssq[2.24398265e-17 2.24398265e-17]

178.FCepsilonRI/k\_f1: 19.2389 FCepsilonRI/k\_f4: 21.0141 FCepsilonRI/k\_f5: 16.9245 FCepsilonRI/k\_f6: 1.96543 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00165854 FCepsilonRI/k\_r4: 0.493027 FCepsilonRI/k\_r6: 5.46755 Ssq[1.15640772e-14 1.15640772e-14]

179.FCepsilonRI/k\_f1: 3.54654 FCepsilonRI/k\_f4: 11.5284 FCepsilonRI/k\_f5: 2.9827 FCepsilonRI/k\_f6: 33.7916 FCepsilonRI/k\_f7: 54.7986 FCepsilonRI/k\_r1: 0.00786154 FCepsilonRI/k\_r4: 15.0399 FCepsilonRI/k\_r6: 5.46755 Ssq[1.26217745e-29 1.26217745e-29]

180.FCepsilonRI/k\_f1: 60.8387 FCepsilonRI/k\_f4: 4.86147 FCepsilonRI/k\_f5: 29.827 FCepsilonRI/k\_f6: 1.90024 FCepsilonRI/k\_f7: 17.2123 FCepsilonRI/k\_r1: 0.0248604 FCepsilonRI/k\_r4: 0.845758 FCepsilonRI/k\_r6: 0.307463 Ssq[2.50960344e-20 2.50960344e-20] 181.FCepsilonRI/k\_f1: 1.12151 FCepsilonRI/k\_f4: 8.86155 FCepsilonRI/k\_f5: 29.827 FCepsilonRI/k\_f6: 1.90024 FCepsilonRI/k\_f7: 17.2123 FCepsilonRI/k\_r1: 0.0248604 FCepsilonRI/k\_r4: 0.845758 FCepsilonRI/k\_r6: 0.307463 Ssq[3.01534572e-17 3.01534572e-17]

182.FCepsilonRI/k\_f1: 1.12151 FCepsilonRI/k\_f4: 4.86147 FCepsilonRI/k\_f5: 29.827 FCepsilonRI/k\_f6: 1.90024 FCepsilonRI/k\_f7: 3.08155 FCepsilonRI/k\_r1: 0.0248604 FCepsilonRI/k\_r4: 0.845758 FCepsilonRI/k\_r6: 0.307463 Ssq[1.61450554e-15 1.61450554e-15]

183.FCepsilonRI/k\_f1: 1.12151 FCepsilonRI/k\_f4: 4.86147 FCepsilonRI/k\_f5: 29.827 FCepsilonRI/k\_f6: 1.90024 FCepsilonRI/k\_f7: 17.2123 FCepsilonRI/k\_r1: 0.00524477 FCepsilonRI/k\_r4: 0.845758 FCepsilonRI/k\_r6: 0.307463 Ssq[3.79866906e-15 3.79866906e-15]

184.FCepsilonRI/k\_f1: 1.12151 FCepsilonRI/k\_f4: 4.86147 FCepsilonRI/k\_f5: 29.827 FCepsilonRI/k\_f6: 1.90024 FCepsilonRI/k\_f7: 17.2123 FCepsilonRI/k\_r1: 0.0248604 FCepsilonRI/k\_r4: 0.027725 FCepsilonRI/k\_r6: 0.307463 Ssq[2.54376432e-19 2.54376432e-19]

185.FCepsilonRI/k\_f1: 1.12151 FCepsilonRI/k\_f4: 4.86147 FCepsilonRI/k\_f5: 29.827 FCepsilonRI/k\_f6: 1.90024 FCepsilonRI/k\_f7: 17.2123 FCepsilonRI/k\_r1: 0.0248604 FCepsilonRI/k\_r4: 0.845758 FCepsilonRI/k\_r6: 12.8495 Ssq[4.59479476e-124.59479476e-12]

186.FCepsilonRI/k\_f1: 1.12151 FCepsilonRI/k\_f4: 4.86147 FCepsilonRI/k\_f5: 29.827 FCepsilonRI/k\_f6: 1.90024 FCepsilonRI/k\_f7: 17.2123 FCepsilonRI/k\_r1: 0.0248604 FCepsilonRI/k\_r4: 0.845758 FCepsilonRI/k\_r6: 0.307463 Ssq[2.71878141e-19 2.71878141e-19]

187.FCepsilonRI/k\_f1: 3.42121 FCepsilonRI/k\_f4: 13.6461 FCepsilonRI/k\_f5: 53.5199 FCepsilonRI/k\_f6: 8.28814 FCepsilonRI/k\_f7: 72.5838 FCepsilonRI/k\_r1: 0.0294935 FCepsilonRI/k\_r4: 0.116915 FCepsilonRI/k\_r6: 54.186 Ssq[4.81868672e-20 4.81868672e-20]

188.FCepsilonRI/k\_f1: 1.99437 FCepsilonRI/k\_f4: 17.7528 FCepsilonRI/k\_f5: 94.3212 FCepsilonRI/k\_f6: 8.28814 FCepsilonRI/k\_f7: 72.5838 FCepsilonRI/k\_r1: 0.00442087 FCepsilonRI/k\_r4: 0.0112784 FCepsilonRI/k\_r6: 23.0565 Ssq[5.56447085e-22 5.56447085e-22]

189.FCepsilonRI/k\_f1: 10.8188 FCepsilonRI/k\_f4: 5.75453 FCepsilonRI/k\_f5: 5.35199 FCepsilonRI/k\_f6: 0.466076 FCepsilonRI/k\_f7: 12.9948 FCepsilonRI/k\_r1: 0.00932667 FCepsilonRI/k\_r4: 0.00657463 FCepsilonRI/k\_r6: 0.00963578 Ssq[9.70114578e-05 9.70114578e-05]

190.FCepsilonRI/k\_f1: 63.0674 FCepsilonRI/k\_f4: 5.75453 FCepsilonRI/k\_f5: 5.35199 FCepsilonRI/k\_f6: 0.466076 FCepsilonRI/k\_f7: 12.9948 FCepsilonRI/k\_r1: 0.00932667 FCepsilonRI/k\_r4: 0.00657463 FCepsilonRI/k\_r6: 0.00963578 Ssq[9.69710319e-05 9.69710319e-05]

191.FCepsilonRI/k\_f1: 10.8188 FCepsilonRI/k\_f4: 42.0986 FCepsilonRI/k\_f5: 5.35199 FCepsilonRI/k\_f6: 0.466076 FCepsilonRI/k\_f7: 12.9948 FCepsilonRI/k\_r1: 0.00932667 FCepsilonRI/k\_r4: 0.00657463 FCepsilonRI/k\_r6: 0.00963578 Ssq[1.83982672e-05 1.83982672e-05]

192.FCepsilonRI/k\_f1: 10.8188 FCepsilonRI/k\_f4: 5.75453 FCepsilonRI/k\_f5: 5.35199 FCepsilonRI/k\_f6: 8.01325 FCepsilonRI/k\_f7: 12.9948 FCepsilonRI/k\_r1: 0.00932667 FCepsilonRI/k\_r4: 0.00657463 FCepsilonRI/k\_r6: 0.00963578 Ssq[4.16935865e-25 4.16935865e-25]

193.FCepsilonRI/k\_f1: 10.8188 FCepsilonRI/k\_f4: 5.75453 FCepsilonRI/k\_f5: 5.35199 FCepsilonRI/k\_f6: 0.466076 FCepsilonRI/k\_f7: 4.08169 FCepsilonRI/k\_r1: 0.00932667 FCepsilonRI/k\_r4: 0.00657463 FCepsilonRI/k\_r6: 0.00963578 Ssq[0.00022661 0.00022661]

194.FCepsilonRI/k\_f1: 10.8188 FCepsilonRI/k\_f4: 5.75453 FCepsilonRI/k\_f5: 5.35199 FCepsilonRI/k\_f6: 0.466076 FCepsilonRI/k\_f7: 12.9948 FCepsilonRI/k\_r1: 0.001398 FCepsilonRI/k\_r4: 0.00657463 FCepsilonRI/k\_r6: 0.00963578 Ssq[9.70043196e-05 9.70043196e-05]

195.FCepsilonRI/k\_f1: 10.8188 FCepsilonRI/k\_f4: 5.75453 FCepsilonRI/k\_f5: 5.35199

FCepsilonRI/k\_f6: 0.466076 FCepsilonRI/k\_f7: 12.9948 FCepsilonRI/k\_r1: 0.00932667 FCepsilonRI/k\_r4: 0.00657463 FCepsilonRI/k\_r6: 1.29656 Ssq[0.00029399 0.00029399]

196.FCepsilonRI/k\_f1: 63.0674 FCepsilonRI/k\_f4: 42.0986 FCepsilonRI/k\_f5: 5.35199 FCepsilonRI/k\_f6: 8.01325 FCepsilonRI/k\_f7: 4.08169 FCepsilonRI/k\_r1: 0.001398 FCepsilonRI/k\_r4: 63.4229 FCepsilonRI/k\_r6: 1.29656 Ssq[3.55111137e-20 3.55111137e-20]

197.FCepsilonRI/k\_f1: 1.12151 FCepsilonRI/k\_f4: 3.4463 FCepsilonRI/k\_f5: 3.23419 FCepsilonRI/k\_f6: 2.1504 FCepsilonRI/k\_f7: 18.8323 FCepsilonRI/k\_r1: 0.0213354 FCepsilonRI/k\_r4: 0.925355 FCepsilonRI/k\_r6: 1.3562 Ssq[5.26062148e-17 5.26062148e-17]

198.FCepsilonRI/k\_f1: 1.24934 FCepsilonRI/k\_f4: 36.7853 FCepsilonRI/k\_f5: 3.23419 FCepsilonRI/k\_f6: 12.5638 FCepsilonRI/k\_f7: 18.8323 FCepsilonRI/k\_r1: 0.0426464 FCepsilonRI/k\_r4: 48.4238 FCepsilonRI/k\_r6: 0.055668 Ssq[1.1359597e-28 1.1359597e-28]

199.FCepsilonRI/k\_f1: 35.4654 FCepsilonRI/k\_f4: 8.17247 FCepsilonRI/k\_f5: 32.3419 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_r1: 0.00674686 FCepsilonRI/k\_r4: 16.4554 FCepsilonRI/k\_r6: 0.0762648 Ssq[1.97215226e-29 1.97215226e-29]

200.FCepsilonRI/k\_f1: 39.5075

1524

FCepsilonRI/k\_f4: 8.17247 FCepsilonRI/k\_f5: 32.3419 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_r1: 0.00674686 FCepsilonRI/k\_r4: 16.4554 FCepsilonRI/k\_r6: 0.0762648 Ssq[7.88860905e-31 7.88860905e-31]

201.FCepsilonRI/k\_f1: 35.4654 FCepsilonRI/k\_f4: 15.5122 FCepsilonRI/k\_f5: 32.3419 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_r1: 0.00674686 FCepsilonRI/k\_r4: 16.4554 FCepsilonRI/k\_r6: 0.0762648 Ssq[2.55590933e-28 2.55590933e-28]

202.FCepsilonRI/k\_f1: 35.4654 FCepsilonRI/k\_f4: 8.17247 FCepsilonRI/k\_f5: 32.0523 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_r1: 0.00674686 FCepsilonRI/k\_r4: 16.4554 FCepsilonRI/k\_r6: 0.0762648 Ssq[3.86541844e-29 3.86541844e-29]

203.FCepsilonRI/k\_f1: 35.4654 FCepsilonRI/k\_f4: 8.17247 FCepsilonRI/k\_f5: 32.3419 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 1.05902 FCepsilonRI/k\_r1: 0.00674686 FCepsilonRI/k\_r4: 16.4554 FCepsilonRI/k\_r6: 0.0762648 Ssq[1.77493704e-28 1.77493704e-28]

204.FCepsilonRI/k\_f1: 35.4654 FCepsilonRI/k\_f4: 8.17247 FCepsilonRI/k\_f5: 32.3419 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_r1: 0.013486 FCepsilonRI/k\_r4: 16.4554 FCepsilonRI/k\_r6: 0.0762648 Ssq[3.15544362e-30 3.15544362e-

#### 30]

205.FCepsilonRI/k\_f1: 35.4654 FCepsilonRI/k\_f4: 8.17247 FCepsilonRI/k\_f5: 32.3419 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_r1: 0.00674686 FCepsilonRI/k\_r4: 0.0086111 FCepsilonRI/k\_r6: 0.0762648 Ssq[7.88860905e-29 7.88860905e-29]

206.FCepsilonRI/k\_f1: 35.4654 FCepsilonRI/k\_f4: 8.17247 FCepsilonRI/k\_f5: 32.3419 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 34.9508 FCepsilonRI/k\_r1: 0.00674686 FCepsilonRI/k\_r4: 16.4554 FCepsilonRI/k\_r6: 0.00313044 Ssq[2.55590933e-28 2.55590933e-28]

207.FCepsilonRI/k\_f1: 39.5075 FCepsilonRI/k\_f4: 15.5122 FCepsilonRI/k\_f5: 32.0523 FCepsilonRI/k\_f6: 38.2401 FCepsilonRI/k\_f7: 1.05902 FCepsilonRI/k\_r1: 0.013486 FCepsilonRI/k\_r4: 0.0086111 FCepsilonRI/k\_r6: 0.00313044 Ssq[6.38977333e-29 6.38977333e-29]

208.FCepsilonRI/k\_f1: 6.30674 FCepsilonRI/k\_f4: 4.2479 FCepsilonRI/k\_f5: 10.1358 FCepsilonRI/k\_f5: 2.97935 FCepsilonRI/k\_f7: 4.46583 FCepsilonRI/k\_r1: 0.00758373 FCepsilonRI/k\_r4: 0.645741 FCepsilonRI/k\_r6: 4.17451 Ssq[2.9945846e-15 2.9945846e-15]

209.FCepsilonRI/k\_f1: 70.2554 FCepsilonRI/k\_f4: 12.585 FCepsilonRI/k\_f5: 10.1358 FCepsilonRI/k\_f6: 2.97935 FCepsilonRI/k\_f7: 4.46583 FCepsilonRI/k\_f1: 0.00758373 FCepsilonRI/k\_r4: 0.645741 FCepsilonRI/k\_r6: 4.17451 Ssq[2.53686513e-13 2.53686513e-13]

210.FCepsilonRI/k\_f1: 70.2554 FCepsilonRI/k\_f4: 4.2479 FCepsilonRI/k\_f5: 10.1358 FCepsilonRI/k\_f6: 2.97935 FCepsilonRI/k\_f7: 4.46583 FCepsilonRI/k\_r1: 0.0119978 FCepsilonRI/k\_r4: 0.645741 FCepsilonRI/k\_r6: 4.17451 Ssq[3.55906457e-143.55906457e-14]

211.FCepsilonRI/k\_f1: 70.2554 FCepsilonRI/k\_f4: 4.2479 FCepsilonRI/k\_f5: 10.1358 FCepsilonRI/k\_f6: 2.97935 FCepsilonRI/k\_f7: 4.46583 FCepsilonRI/k\_r1: 0.00758373 FCepsilonRI/k\_r4: 0.645741 FCepsilonRI/k\_r6: 0.0180852 Ssq[3.02964301e-21 3.02964301e-21]

212.FCepsilonRI/k\_f1: 70.2554 FCepsilonRI/k\_f4: 4.2479 FCepsilonRI/k\_f5: 10.1358 FCepsilonRI/k\_f5: 2.97935 FCepsilonRI/k\_f7: 4.46583 FCepsilonRI/k\_r1: 0.00758373 FCepsilonRI/k\_r4: 0.645741 FCepsilonRI/k\_r6: 4.17451 Ssq[1.89072327e-15 1.89072327e-15]

213.FCepsilonRI/k\_f1: 63.0674 FCepsilonRI/k\_f4: 70.7707 FCepsilonRI/k\_f5: 1.02274 FCepsilonRI/k\_f6: 9.06817 FCepsilonRI/k\_f7: 8.28814 FCepsilonRI/k\_r1: 0.00119978 FCepsilonRI/k\_r4: 0.219436 FCepsilonRI/k\_r6: 5.71905 Ssq[2.1825888e-25 2.1825888e-25]

214.FCepsilonRI/k\_f1: 7.02554 FCepsilonRI/k\_f4: 70.7707 FCepsilonRI/k\_f5: 1.02274 FCepsilonRI/k\_f6: 9.06817 FCepsilonRI/k\_f7: 8.28814 FCepsilonRI/k\_r1: 0.00119978 FCepsilonRI/k\_r4: 0.219436 FCepsilonRI/k\_r6: 5.71905 Ssq[1.01290234e-20 1.01290234e-20]

215.FCepsilonRI/k\_f1: 63.0674 FCepsilonRI/k\_f4: 23.8877 FCepsilonRI/k\_f5: 1.02274 FCepsilonRI/k\_f5: 9.06817 FCepsilonRI/k\_f7: 8.28814 FCepsilonRI/k\_r1: 0.00119978 FCepsilonRI/k\_r4: 0.219436 FCepsilonRI/k\_r6: 5.71905 Ssq[6.41393527e-23 6.41393527e-23]

216.FCepsilonRI/k\_f1: 63.0674 FCepsilonRI/k\_f4: 70.7707 FCepsilonRI/k\_f5: 1.02274 FCepsilonRI/k\_f6: 9.06817 FCepsilonRI/k\_f7: 8.28814 FCepsilonRI/k\_r1: 0.0758373 FCepsilonRI/k\_r4: 0.219436 FCepsilonRI/k\_r6: 5.71905 Ssq[2.10853842e-21 2.10853842e-21]

217.FCepsilonRI/k\_f1: 63.0674 FCepsilonRI/k\_f4: 70.7707 FCepsilonRI/k\_f5: 1.02274 FCepsilonRI/k\_f6: 9.06817 FCepsilonRI/k\_f7: 8.28814 FCepsilonRI/k\_r1: 0.00119978 FCepsilonRI/k\_r4: 0.00204201 FCepsilonRI/k\_r6: 5.71905 Ssq[3.33710223e-19 3.33710223e-19]

218.FCepsilonRI/k\_f1: 63.0674 FCepsilonRI/k\_f4: 70.7707 FCepsilonRI/k\_f5: 1.02274 FCepsilonRI/k\_f6: 9.06817 FCepsilonRI/k\_f7: 8.28814 FCepsilonRI/k\_r1: 0.00119978 FCepsilonRI/k\_r4: 0.219436 FCepsilonRI/k\_r6: 0.013201 Ssq[1.1359597e-28 1.1359597e-28] 219.FCepsilonRI/k\_f1: 93.6871 FCepsilonRI/k\_f4: 70.2949 FCepsilonRI/k\_f5: 1.80243 FCepsilonRI/k\_f6: 6.11816 FCepsilonRI/k\_f7: 4.03605 FCepsilonRI/k\_r1: 0.00179839 FCepsilonRI/k\_r4: 0.0176831 FCepsilonRI/k\_r6: 2.03285 Ssq[1.37174234e-25 1.37174234e-25]

220.FCepsilonRI/k\_f1: 8.41017 FCepsilonRI/k\_f4: 24.0494 FCepsilonRI/k\_f5: 1.81872 FCepsilonRI/k\_f6: 0.34405 FCepsilonRI/k\_f7: 71.7723 FCepsilonRI/k\_r1: 0.0284513 FCepsilonRI/k\_r4: 0.00600909 FCepsilonRI/k\_r6: 0.00208845 Ssq[0.00310188 0.00310188]

221.FCepsilonRI/k\_f1: 8.41017 FCepsilonRI/k\_f4: 29.6431 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f5: 0.34405 FCepsilonRI/k\_f7: 38.6725 FCepsilonRI/k\_r1: 0.0568699 FCepsilonRI/k\_r4: 0.314455 FCepsilonRI/k\_r6: 0.114316 Ssq[0.00044038 0.00044038]

222.FCepsilonRI/k\_f1: 29.6265 FCepsilonRI/k\_f4: 24.0494 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f5: 0.34405 FCepsilonRI/k\_f7: 38.6725 FCepsilonRI/k\_r1: 0.0568699 FCepsilonRI/k\_r4: 0.314455 FCepsilonRI/k\_r6: 0.114316 Ssq[0.00047591 0.00047591]

223.FCepsilonRI/k\_f1: 29.6265 FCepsilonRI/k\_f4: 29.6431 FCepsilonRI/k\_f5: 1.81872 FCepsilonRI/k\_f5: 0.34405 FCepsilonRI/k\_f7: 38.6725 FCepsilonRI/k\_r1: 0.0568699 FCepsilonRI/k\_r4: 0.314455 FCepsilonRI/k\_r6: 0.114316 Ssq[0.00329041 0.00329041]

224.FCepsilonRI/k\_f1: 29.6265

FCepsilonRI/k\_f4: 29.6431 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f6: 0.34405 FCepsilonRI/k\_f7: 71.7723 FCepsilonRI/k\_r1: 0.0568699 FCepsilonRI/k\_r4: 0.314455 FCepsilonRI/k\_r6: 0.114316 Ssq[0.00041535] 0.00041535]

225.FCepsilonRI/k\_f1: 29.6265 FCepsilonRI/k\_f4: 29.6431 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f6: 0.34405 FCepsilonRI/k\_f7: 38.6725 FCepsilonRI/k\_r1: 0.0284513 FCepsilonRI/k\_r4: 0.314455 FCepsilonRI/k\_r6: 0.114316 Ssq[0.00044001 0.00044001] 226.FCepsilonRI/k\_f1: 29.6265 FCepsilonRI/k\_f4: 29.6431 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f6: 0.34405 FCepsilonRI/k\_f7: 38.6725 FCepsilonRI/k\_r1: 0.0568699 FCepsilonRI/k\_r4: 0.00600909 FCepsilonRI/k\_r6: 0.114316 Ssq[0.00043743 0.00043743] 227.FCepsilonRI/k\_f1: 29.6265 FCepsilonRI/k\_f4: 29.6431

FCepsilonRI/k\_f4: 29.6431 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f6: 0.34405 FCepsilonRI/k\_f7: 38.6725 FCepsilonRI/k\_r1: 0.0568699 FCepsilonRI/k\_r4: 0.314455 FCepsilonRI/k\_r6: 0.00208845 Ssq[0.00042624 0.00042624] 228.FCepsilonRI/k\_f1: 29.6265 FCepsilonRI/k\_f4: 29.6431 FCepsilonRI/k\_f5: 18.0243 FCepsilonRI/k\_f5: 0.34405 FCepsilonRI/k\_f7: 38.6725 FCepsilonRI/k\_r1: 0.0568699 FCepsilonRI/k\_r4: 0.314455 FCepsilonRI/k\_r6: 0.114316 Ssq[0.00044002 0.00044002]