# Chronic Myeloid Leukaemia Role of Transplantation

Jane Apperley



- Male aged 30 presents with CML in 1<sup>st</sup> chronic phase
- Intermediate Sokal score
- HLA identical brother
- EBMT (Gratwohl) score = 1 (affected by age, disease phase, type of donor, donor-recipient genders, time from diagnosis)

# 1997

### **Outcome of allo-SCT for CML by EBMT risk score (2011)**



Courtesy of R Brand, Oct 2011



- Male aged 30 presents with CML in CP1
- Intermediate Sokal score
- HLA identical brother
- Allo-SCT or imatinib?

## **Transplant activity since 1980 at Imperial College**





# **IRIS 8-year update: overall survival**



Deininger M, et al. Blood. 2009;114(22):462. Abstract # 1126.





- Male aged 30 presents with CML CP1: intermediate Sokal score: HLA-identical brother
- Starts imatinib 400mg daily, tolerated well
- At 12 months 100% Ph-neg, RT-PCR 1.8%
- At 18 months, RT-PCR 11%
- Bone marrow confirms loss of complete cytogenetic response
- Patient insists that he is compliant





- Male aged 30 presents with CML CP1: intermediate Sokal score: HLA-identical brother
- Starts imatinib 400mg daily
- Achieves complete cytogenetic response at 12 months but loses this at 18 months
- Allo-SCT or a 2<sup>nd</sup> generationTKI (2GTKI), i.e. dasatinib or nilotinib?

### 2101 CML-CP: nilotinib for imatinib failure (n=321)



Kantarjian et al. Haematologica (14th Annual Congress of the EHA). 2009;94(s1): Abstract 0627.

# Failure after imatinib

**Responders to 2GTKI are highly predictable** 

- Sokal score at diagnosis
- Degree of cytogenetic response to imatinib
- Presence or absence of neutropenia on imatinib
- Short-term (3 months) response to 2GTKI

# Case History: 2007

- Male aged 30 presents with CML CP1: intermediate Sokal score: HLA-identical brother
- Starts imatinib 400mg daily
- Achieves CCyR at 12 months but loses this at 18 months
- 2GTKI for 6 months, achieves CCyR but not major molecular response, drug is poorly tolerated
- Allo-SCT, stay on current therapy or change to a 3GTKI

# **Outcome of 2GTKI in 110 patients**



Time (years) from the onset of second line therapy

Milojkovic et al, Blood 2011

# Ponatinib Phase 2 Study - PACE Response by Number of Prior Approved TKIs CP-CML

📕 1 TKI 📕 2 TKIS 📕 3 TKIS



TKIs = imatinib, dasatinib, nilotinib

\*Includes 3 CP-CML patients who were non-cohort assigned (post-imatinib, non-T315I), but treated



- Male aged 30 presents with CML in CP1
- Intermediate Sokal score
- HLA identical brother
- Imatinib or 2GTKI or clinical trial?

# **Outcome of frontline imatinib**



# **Outcome of frontline imatinib**



# **Outcome of frontline 2GTKI**

![](_page_16_Figure_2.jpeg)

# **Outcome of frontline 2GTKI**

![](_page_17_Figure_2.jpeg)

### **Outcome of frontline 3GTKI?**

![](_page_18_Figure_2.jpeg)

10-15% require alternative therapy

### **Outcome of allo-SCT for CML by EBMT risk score (2011)**

![](_page_19_Figure_2.jpeg)

Courtesy of R Brand, Oct 2011

#### **Outcome of transplantation for advanced phase disease**

![](_page_20_Figure_2.jpeg)

![](_page_21_Picture_1.jpeg)

Mary Alikian, Alex Bazeos, Marco Bua, Gordon Cook, Letizia Foroni, Gareth Gerrard, John Goldman, Ailsa Holroyd, David Marin, Dragana Milojkovic, George Nteliopoulos, Christos Paliompeis, Alistair Reid, Richard Szydlo, BSBMT teams and colleagues at Ariad, BMS, Novartis and Pfizer

![](_page_21_Picture_3.jpeg)

**Beating Blood Cancers** 

![](_page_21_Picture_5.jpeg)

![](_page_21_Picture_6.jpeg)