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PRESENTATIONS OF "BIPOLAR SPECTRUM": IMPLICATIONS FOR REFORM IN CLINICAL PRACTICE

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### WHAT IS THE QUESTION?

Does the structure of reality of mood disorders allow cleavage of discrete diagnostic categories?

### or

Would mood disorders be considered as a continuous spectrum?

### Dichotomous vs Unitary



 Based on dichotomous positions proposed by Neele (1949), Leonhard (1958), Angst (1966/1973), Perris (1966), and Winokur et al. (1969), in the ICD-9 and 10; and DSM-III, and IV, bipolar (BP) and major depressive disorders of the unipolar type (MDD, UP) were distinct categories.

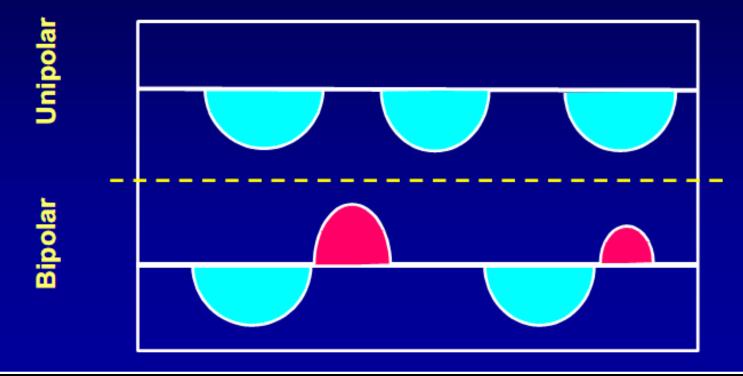
This is contrary to Kraepelin's (1899/1921) earlier unitary concept of manic-depression, which included manic, circular, as well as recurrent depressive conditions.



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Manic-Depressive Psychosis A Unitary Entity (Kraepelin, 1899)

Later dichotomized into unipolar versus bipolar (Angst 1966; Perris 1966; Winokur & Clayton 1969)



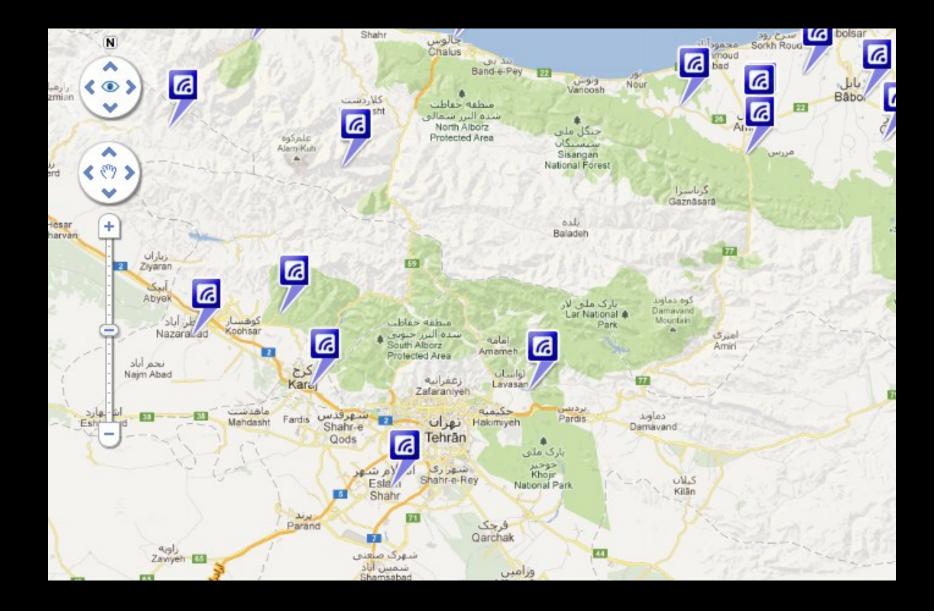




- Are the borders real? Or they are imaginary lines on sands!
- Do you think what are borders?
- What is the advantage of a border?

### Let's Have a Trip!





















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- A disorder is determine by:
  - Multiple, well defined symptoms,
  - Distinct, prolonged duration,
  - Critical, disturbed function





Akiskal HS, Djenderedjian AH, Rosenthal RH, Khani MK.1977. Cyclothymic disorder: validating criteria for inclusion in the bipolar affective group.

Am. J. Psychiatry. 134, 1227–1233.

 DSM-IV: 2 years of numerous periods with hypomanic symptoms and numerous periods with depressive symptoms that do not meet criteria for a major depressive episode,





- Iow self-esteem or self-confidence, or feelings of inadequacy
- feelings of pessimism, despair, or hopelessness
- generalized loss of interest or pleasure
- social withdrawal
- chronic fatigue or tiredness
- feelings of guilt, brooding about the past
- subjective feelings of irritability or excessive anger
- decreased activity, effectiveness, or productivity
- difficulty in thinking, reflected by poor concentration, poor memory, or indecisiveness



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- Impulsive
- Extremely outgoing
- Competitive
- Have a great deal of energy
- Aspiring
- Enthusiastic
- High productivity





 Akiskal HS. 1981. Subaffective disorders: dysthymic, cyclothymic and bipolar II disorders in the 'borderline' realm.

Psychiatr Clin NorthAm 4,25–46.

Akiskal HS. 1983. The bipolar spectrum: New concepts in classification and diagnosis. In: Grinspoon L (Ed.), Psychiatry Update: The American Psychiatric Association Annual Review, vol.2. American Psychiatric Press, Washington, DC, pp.271–292.



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 Akiskal HS, Mallya G. 1987. Criteria for the 'soft' bipolar spectrum: treatment implications.
 Psychopharmacol Bull. 23, 68–73.

 Akiskal HS, Cassano GB, Musetti L, Perugi G, Tundo A, Mignani V. 1989. Psychopathology, temperament and past course in primary major depressions: Review of evidence for a bipolar spectrum.

Psychopathology 22,268–277.



 Akiskal HS. 1992. Delineating irritable-choleric and hyperthymic temperaments as variants of cyclothymia.

J Person Disord. 6,326–342.

 Akiskal HS. Akiskal K. 1992. Cyclothymic, hyperthymic and depressive temperaments as subaffective variants of mood disorders. In: Tasman A, Riba MB (Eds.), American Psychiatric Association Review. American Psychiatric Press, Washington, DC,. 43–62.



# Attributes, Assets, and Liabilities of Temperaments

Depressive	Hyperthymic
Gloomy, incapable of fun, complaining	Cheerful and exuberant
Humorless	Articulate and jocular
Pessimistic and given to brooding	Overoptimistic and carefree
Guilt-prone, low self-esteem, and preoccupied with inadequacy or failure	Overconfident, self-assured, boastful, and grandiose
Introverted with restricted social life	Extroverted and people seeking
Sluggish, living a life out of action	High energy level, full of plans
Few but constant interests	Versatile with broad interests
Passive	Overinvolved and meddlesome
Reliable, dependable, and devoted	Uninhibited and stimulus seeking



Akiskal HS. 1996. The prevalent clinical spectrum of bipolar disorders: beyond DSM-IV.

J Clin psycho-pharmacol. 16 (Suppl1),4s–14s.

 Akiskal HS, Pinto OC. 1999. The evolving bipolar spectrum. Prototypes I, II, II, and IV.
 Psychiatr Clin NorthAm 22,517–534.

Angst J, Gamma A. 2002. A new bipolar spectrum concept: a brief review.

Bipolar Disord. 4 (Suppl 1), 11–14.



Journal of Affective Disorders 107 (2008) 307-315



www.elsevier.com/locate/jad

Preliminary communication

#### A case series on the hypothesized connection between dementia and bipolar spectrum disorders: Bipolar type VI?

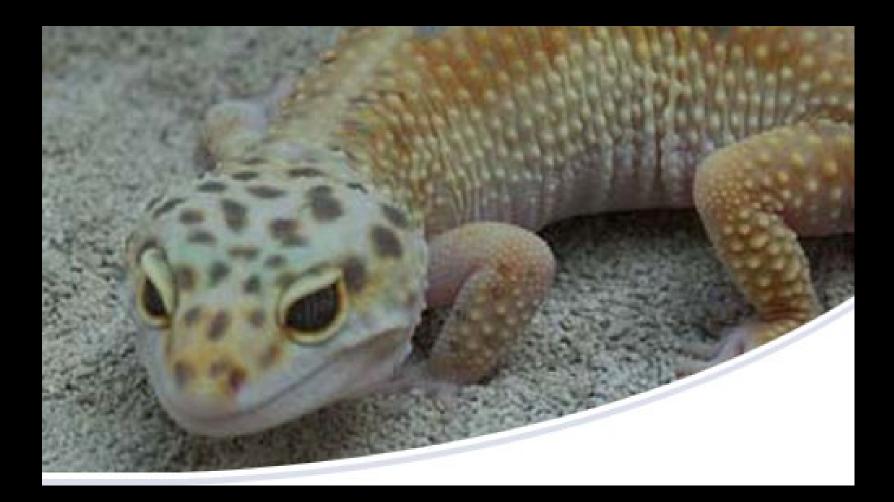
Bernardo Ng<sup>a,b,c,\*</sup>, Alvaro Camacho<sup>a,b,c</sup>, Diogo R. Lara<sup>d</sup>, Miriam G. Brunstein<sup>e</sup>, Olavo C. Pinto<sup>f</sup>, Hagop S. Akiskal<sup>a,b</sup>

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> > Received 4 January 2007; accepted 28 August 2007 Available online 21 September 2007



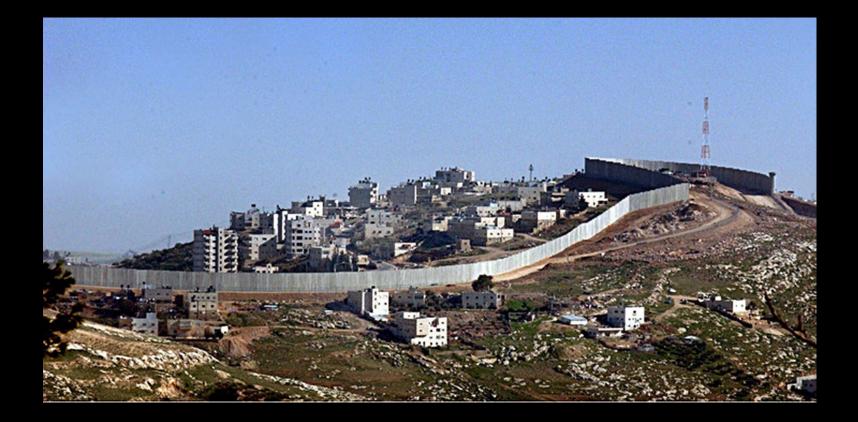




### **History Hates Borders!**



### Shame on Border Makers!



# The Final Destiny!





JOURNAL



Journal of Affective Disorders 54 (1999) 319-328

Special article

# The bipolar spectrum: a clinical reality in search of diagnostic criteria and an assessment methodology

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Received 3 May 1998; accepted 7 October 1998



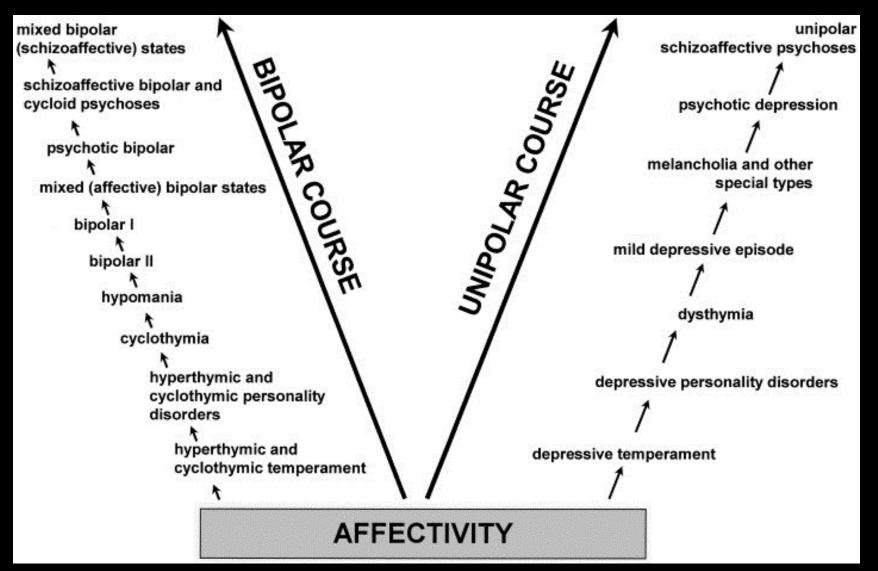


- From unipolar depression to bipolar I,
- From infrequent bipolar episodes to continuous cycling (cycle rate as a dimensional concept),
- Continuum of mixed states, from those meeting full manic and depressive criteria, to depression which appears purely unipolar but a few or even a single manic symptom,
- From bipolar disorder to schizophrenia,



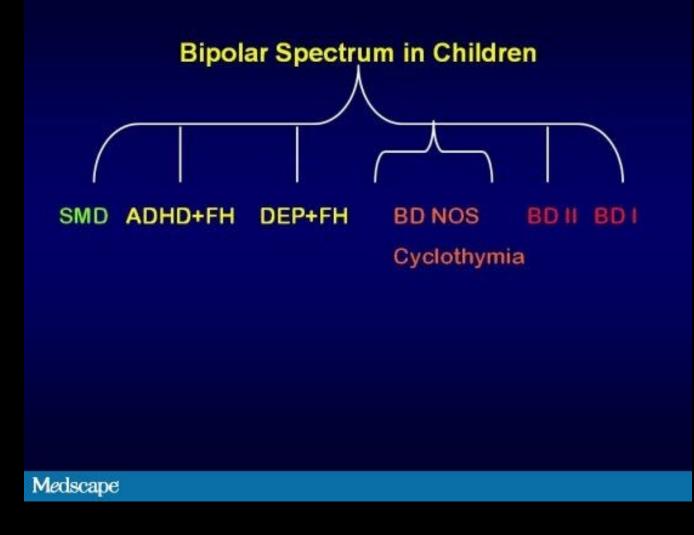
- From bipolar disorder to borderline personality disorder,
- From bipolar spectrum to ADHD,
- From bipolar spectrum to impulse control disorders,
- Apparent continuum of symptom presentation from normal to pathologic,







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#### Comorbidities of impulse control disorders

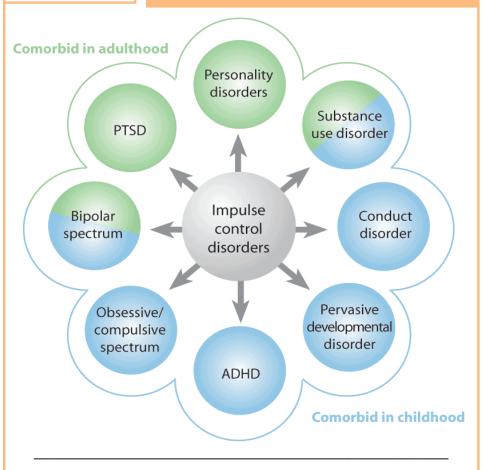
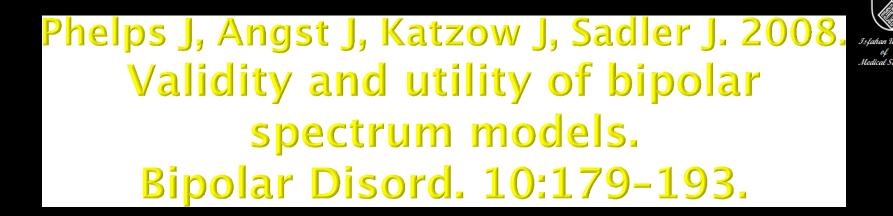


Figure 3

ADHD, attention-deficit/hyperactivity disorder; PTSD, posttraumatic stress disorder.



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Criteria

'Intuitive' selection then prospective evaluation

Literature review of the 'spectrum of mania'

Bipolar markers grouped in five dimensions

List of non-manic bipolar markers; DSM-like schema

Retrospective subgrouping

Narrative, case illustrations

Epidemiology based

Table 1. Diagnostic schema for a bipolar spectrum

Year

1977

1978

1981

1999

2002

2003

2004

System

Categories I-VI

Cyclothymia-bipolar continuum

Add minor bipolar disorders

100-point 'Bipolarity Index'

Bipolar subtypes along a spectrum

Categories I-IV including 11/2, 21/2, 31/2

New category: 'bipolar spectrum disorder'

Authors

Angst (88)

Klerman (30)

Akiskal et al. (29)

Akiskal and Pinto (31)

Ghaemi et al. (50)

Angst et al. (46)

Sachs (67)





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Psychotic (mood congruent or mood incongruent)	D	Dm	MD	Md	М	
Non-psychotic	D	Dm	MD	Md	м	
Subthreshold	d		md		m	
'Normal'	dsx		mdsx		msx	



- The problem with lowering cut-offs, is that setting the bar for a diagnosis anywhere below full mania makes the condition difficult to define operationally.
- Decreased Inter-rater reliability (especially between incompetence raters),
- Specificity immolate for higher sensitivity,

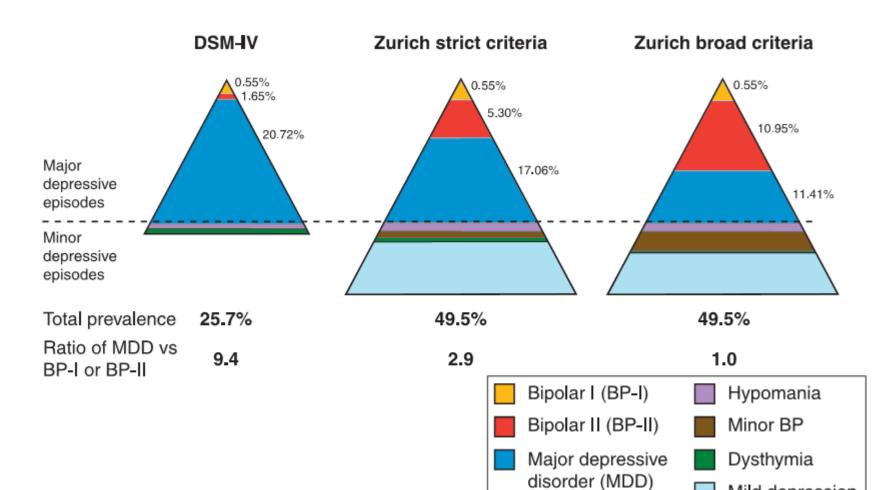
Higher prevalence rate,



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Mild depression

# Epidemiology: The Trick!



Bipolarity independent of hypomania or mania (Bipolar markers)



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The criteria of Ghaemi et al. (2002) for the BP spectrum disorders restricted the concept of BP spectrum to what is beyond DSM-IV BP-II.

## **MDE plus indicators of bipolarity:**

- Bipolar family history,
- Early age at onset,
- High depressive recurrence,

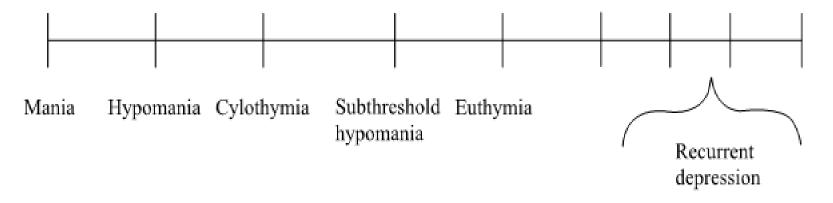


- Antidepressant-associated hypomania,
- Atypical features,
- Postpartum onset
- Hyperthymic temperament





Mild Moderate Severe Psychotic





- Oops! What was happened? We have a new category!
- Where is the spectrum in this purposed category!?
- We have a new box for putting those patients who do not settle in previous boxes!
- Up to what number we can increase the boxes?







#### Table 13.7–34. Evolving Spectrum of Bipolar Disorders

Bipolar <sup>1</sup>/<sub>2</sub>: Schizobipolar disorder
Bipolar I: Core manic-depressive illness
Bipolar I<sup>1</sup>/<sub>2</sub>: Depression with protracted hypomania
Bipolar II: Depression with discrete spontaneous hypomanic episodes
Bipolar II<sup>1</sup>/<sub>2</sub>: Depression superimposed on cyclothymic temperament
Bipolar III: Depression plus *induced* hypomania (i.e., hypomania occurring solely in association with antidepressant or other somatic treatment)
Bipolar III<sup>1</sup>/<sub>2</sub>: Prominent mood swings occurring in the context of

Bipolar III<sup>1</sup>/<sub>2</sub>: Prominent mood swings occurring in the context of substance or alcohol use or abuse

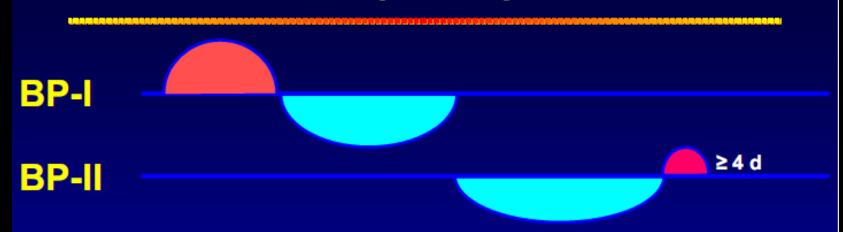
Bipolar IV: Depression superimposed on a hyperthymic temperament

Adapted from Akiskal HS, Pinto O: The evolving bipolar spectrum: Prototypes I, II, III, IV. *Psychiatr Clin North Am.* 1999;22:517.



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#### **Akiskal Bipolar Spectrum**



#### 

Cyclothymic Depression

Antidepressants

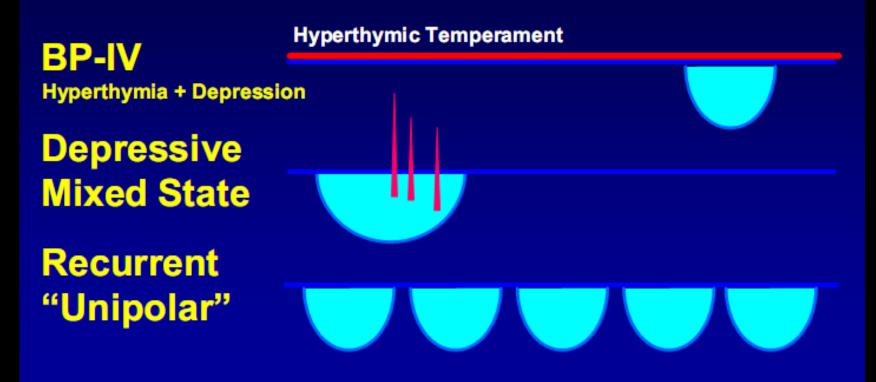
BP-III Pharmacologic Hypomania

Akiskal HS, Pinto O. Psychiatr Clin North Am 1999;22:517-34.



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### Other Proposed Akiskal Bipolar Spectrum Subtypes



Akiskal, et al. J Affect Disord 2000;59(Suppl 1):S5-S30.





- Allows for the illness to exist as a matter of degree, and places patients on a presumed continuum of bipolarity.
- This instrument is currently in use as part of the Massachusetts General Hospital Bipolar Clinics Affective Disorders Evaluation.



- In the Index, bipolar markers are organized in five dimensions, each worth 20 points, for a 100-point total scale:
  - 1) Episode characteristics
  - 2) Age of onset
  - 3) Course of illness/associated features
  - 4) Response to treatment
  - 5) Family history



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الهى قطر، دانش که بخشیدی زپیش متصل گردان به در پاهای خویش